PCT

(22) International Filing Date:

60/096,116

WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁶ : C12N 15/12, C07K 14/47, 16/18, C12Q 1/68		Δ2	(11) International Publication Number	: WO 99/31236
			(43) International Publication Date:	24 June 1999 (24.06.99)
(21) International Application Number:	PCT/IB	98/021:	(81) Designated States: AL, AM, A	r, AU, AZ, BA, BB, BG, BR,

US

17 December 1998 (17.12.98)

(30) Priority Data:
60/069,957
60/074,121
60/081,563
17 December 1997 (17.12.97)
9 February 1998 (09.02.98)
US
US

10 August 1998 (10.08.98)

(71) Applicant (for all designated States except US): GENSET [FR/FR]; 24, rue Royale, F-75008 Paris (FR).

(72) Inventors; and
(75) Inventors/Applicants (for US only): BOUGUELERET, Lydie [FR/FR]; 108, avenue Victor Hugo, F-92170 Vanves (FR). DUCLERT, Aymeric [FR/FR]; 6 ter, rue Victorine, F-94100 Saint-Maur (FR). DUMAS MILNE EDWARDS, Jean-Baptiste [FR/FR]; 8, rue Grégoire de Tours, F-75006 Paris (FR).

(74) Agents: MARTIN, Jean-Jacques et al.; Cabinet Regimbeau, 26, avenue Kléber, F-75116 Paris (FR).

(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Published

Without international search report and to be republished upon receipt of that report.

(54) Title: EXTENDED cDNAs FOR SECRETED PROTEINS

(57) Abstract

The sequences of extended cDNAs encoding secreted proteins are disclosed. The extended cDNAs can be used to express secreted proteins or portions thereof or to obtain antibodies capable of specifically binding to the secreted proteins. The extended cDNAs may also be used in diagnostic, forensic, gene therapy, and chromosome mapping procedures. The extended cDNAs may also be used to design expression vectors and secretion vectors.

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Amenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT		FR	France	LU	Luxembourg	SN	Senegal
	Austria	GA	Gabon	LV	Latvia	SZ	Swaziland
ΑU	Australia	GB	United Kingdom	MC	Monaco	TD	Chad
AZ	Azerbaijan	GE	Georgia	MD	Republic of Moldova	TG	Togo
BA	Bosnia and Herzegovina		-	MG	Madagascar	TJ	Tajikistan
BB	Barbados	GH	Ghana	MK MK	The former Yugoslav	TM	Turkmenistan
BE	Belgium	GN	Guinea	WIR	Republic of Macedonia	TR	Turkey
BF	Burkina Faso	GR	Greece		-	TT	Trinidad and Tobago
BG	Bulgaria	HU	Hungary	ML	Mali	UA	Ukraine
BJ	Benin	IE	Ireland	MN	Mongolia		
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
CH	Switzerland	KG	Kyrgyzstan	NO	Norway	zw	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's	NZ	New Zealand		
CM	Cameroon		Republic of Korea	PL	Poland		
CN	China	KR	Republic of Korea	PT	Portugal		
		KZ	Kazakstan	RO	Romania		
CU	Cuba			RU	Russian Federation		
CZ	Czech Republic	LC	Saint Lucia				
DE	Germany	L,I	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		

WO 99/31236 PCT/IB98/02122

EXTENDED cDNAS for secreted proteins

The present application relates to extended cDNAs which were disclosed in several United States Provisional Patent Applications. Table I lists the SEQ ID Nos. of the extended cDNAs in the present application, the SEQ ID Nos. of the identical or nearly identical extended cDNAs in the provisional applications, and the identities of the provisional applications in which the extended cDNAs were disclosed.

Background of the Invention

The estimated 50,000-100,000 genes scattered along the human chromosomes offer tremendous promise for the understanding, diagnosis, and treatment of human diseases. In addition, probes capable of specifically hybridizing to loci distributed throughout the human genome find applications in the construction of high resolution chromosome maps and in the identification of individuals.

In the past, the characterization of even a single human gene was a painstaking process, requiring years of effort. Recent developments in the areas of cloning vectors, DNA sequencing, and computer technology have merged to greatly accelerate the rate at which human genes can be isolated, sequenced, mapped, and characterized. Cloning vectors such as yeast artificial chromosomes (YACs) and bacterial artificial chromosomes (BACs) are able to accept DNA inserts ranging from 300 to 1000 kilobases (kb) or 100-400 kb in length respectively, thereby facilitating the manipulation and ordering of DNA sequences distributed over great distances on the human chromosomes. Automated DNA sequencing machines permit the rapid sequencing of human genes. Bioinformatics software enables the comparison of nucleic acid and protein sequences, thereby assisting in the characterization of human gene products.

Currently, two different approaches are being pursued for identifying and characterizing the genes distributed
20 along the human genome. In one approach, large fragments of genomic DNA are isolated, cloned, and sequenced.

Potential open reading frames in these genomic sequences are identified using bio-informatics software. However, this approach entails sequencing large stretches of human DNA which do not encode proteins in order to find the protein encoding sequences scattered throughout the genome. In addition to requiring extensive sequencing, the bio-informatics software may mischaracterize the genomic sequences obtained. Thus, the software may produce-false positives in which non-coding DNA is mischaracterized as coding DNA or false negatives in which coding DNA is mislabeled as non-coding DNA.

An alternative approach takes a more direct route to identifying and characterizing human genes. In this approach, complementary DNAs (cDNAs) are synthesized from isolated messenger RNAs (mRNAs) which encode human proteins. Using this approach, sequencing is only performed on DNA which is derived from protein coding portions of the genome. Often, only short stretches of the cDNAs are sequenced to obtain sequences called expressed sequence tags (ESTs). The ESTs may then be used to isolate or purify extended cDNAs which include sequences adjacent to the EST sequences. The extended cDNAs may contain all of the sequence of the EST which was used to obtain them or only a portion of the sequence of the EST which was used to obtain them. In addition, the extended cDNAs may contain the full coding sequence of the gene from which the EST was derived or, alternatively, the extended cDNAs may include

portions of the coding sequence of the gene from which the EST was derived. It will be appreciated that there may be several extended cDNAs which include the EST sequence as a result of alternate splicing or the activity of alternative promoters.

In the past, the short EST sequences which could be used to isolate or purify extended cDNAs were often 5 obtained from oligo-dT primed cDNA libraries. Accordingly, they mainly corresponded to the 3' untranslated region of the mRNA. In part, the prevalence of EST sequences derived from the 3' end of the mRNA is a result of the fact that typical techniques for obtaining cDNAs, are not well suited for isolating cDNA sequences derived from the 5' ends of mRNAs. (Adams et al., Nature 377:174, 1996, Hillier et al., Genome Res. 6:807-828, 1996).

In addition, in those reported instances where longer cDNA sequences have been obtained, the reported 10 sequences typically correspond to coding sequences and do not include the full 5' untranslated region of the mRNA from which the cDNA is derived. Such incomplete sequences may not include the first exon of the mRNA, particularly in situations where the first exon is short. Furthermore, they may not include some exons, often short ones, which are located upstream of splicing sites. Thus, there is a need to obtain sequences derived from the 5' ends of mRNAs which can be used to obtain extended cDNAs which may include the 5' sequences contained in the 5' ESTs.

While many sequences derived from human chromosomes have practical applications, approaches based on the identification and characterization of those chromosomal sequences which encode a protein product are particularly relevant to diagnostic and therapeutic uses. Of the 50,000-100,000 protein coding genes, those genes encoding proteins which are secreted from the cell in which they are synthesized, as well as the secreted proteins themselves, are particularly valuable as potential therapeutic agents. Such proteins are often involved in cell to cell communication and 20 may be responsible for producing a clinically relevant response in their target cells.

In fact, several secretory proteins, including tissue plasminogen activator, G-CSF, GM-CSF, erythropoietin, human growth hormone, insulin, interferon- α , interferon- β , interferon- γ , and interleukin-2, are currently in clinical use. These proteins are used to treat a wide range of conditions, including acute myocardial infarction, acute ischemic stroke, anemia, diabetes, growth hormone deficiency, hepatitis, kidney carcinoma, chemotherapy induced neutropenia and 25 multiple sclerosis. For these reasons, extended cDNAs encoding secreted proteins or portions thereof represent a particularly valuable source of therapeutic agents. Thus, there is a need for the identification and characterization of secreted proteins and the nucleic acids encoding them.

In addition to being therapeutically useful themselves, secretory proteins include short peptides, called signal peptides, at their amino termini which direct their secretion. These signal peptides are encoded by the signal sequences 30 located at the 5' ends of the coding sequences of genes encoding secreted proteins. Because these signal peptides will direct the extracellular secretion of any protein to which they are operably linked, the signal sequences may be exploited to direct the efficient secretion of any protein by operably linking the signal sequences to a gene encoding the protein for which secretion is desired. This may prove beneficial in gene therapy strategies in which it is desired to deliver a particular gene product to cells other than the cell in which it is produced. Signal sequences encoding signal peptides

also find application in simplifying protein purification techniques. In such applications, the extracellular secretion of the desired protein greatly facilitates purification by reducing the number of undesired proteins from which the desired protein must be selected. Thus, there exists a need to identify and characterize the 5' portions of the genes for secretory proteins which encode signal peptides.

Public information on the number of human genes for which the promoters and upstream regulatory regions have been identified and characterized is quite limited. In part, this may be due to the difficulty of isolating such regulatory sequences. Upstream regulatory sequences such as transcription factor binding sites are typically too short to be utilized as probes for isolating promoters from human genomic libraries. Recently, some approaches have been developed to isolate human promoters. One of them consists of making a CpG island library (Cross, S.H. et al., 10 Purification of CpG Islands using a Methylated DNA Binding Column, Nature Genetics 6: 236-244 (1994)). The second consists of isolating human genomic DNA sequences containing Spel binding sites by the use of Spel binding protein. (Mortlock et al., Genome Res. 6:327-335, 1996). Both of these approaches have their limits due to a lack of specificity or of comprehensiveness.

5' ESTs and extended cDNAs obtainable therefrom may be used to efficiently identify and isolate upstream 15 regulatory regions which control the location, developmental stage, rate, and quantity of protein synthesis, as well as the stability of the mRNA. (Theil et al., BioFactors 4:87-93, (1993). Once identified and characterized, these regulatory regions may be utilized in gene therapy or protein purification schemes to obtain the desired amount and locations of protein synthesis or to inhibit, reduce, or prevent the synthesis of undesirable gene products.

In addition, ESTs containing the 5' ends of secretory protein genes or extended cDNAs which include 20 sequences adjacent to the sequences of the ESTs may include sequences useful as probes for chromosome mapping and the identification of individuals. Thus, there is a need to identify and characterize the sequences upstream of the 5' coding sequences of genes encoding secretory proteins.

Summary of the Invention

The present invention relates to purified, isolated, or recombinant extended cDNAs which encode secreted 25 proteins or fragments thereof. Preferably, the purified, isolated or recombinant cDNAs contain the entire open reading frame of their corresponding mRNAs, including a start codon and a stop codon. For example, the extended cDNAs may include nucleic acids encoding the signal peptide as well as the mature protein. Alternatively, the extended cDNAs may contain a fragment of the open reading frame. In some embodiments, the fragment may encode only the sequence of the mature protein. Alternatively, the fragment may encode only a portion of the mature protein. A further aspect of the 30 present invention is a nucleic acid which encodes the signal peptide of a secreted protein.

The present extended cDNAs were obtained using ESTs which include sequences derived from the authentic 5' ends of their corresponding mRNAs. As used herein the terms "EST" or "5' EST" refer to the short cDNAs which were used to obtain the extended cDNAs of the present invention. As used herein, the term "extended cDNA" refers to the cDNAs which include sequences adjacent to the 5' EST used to obtain them. The extended cDNAs may contain all or a

portion of the sequence of the EST which was used to obtain them. The term "corresponding mRNA" refers to the mRNA which was the template for the cDNA synthesis which produced the 5' EST. As used herein, the term "purified" does not require absolute purity; rather, it is intended as a relative definition. Individual extended cDNA clones isolated from a cDNA library have been conventionally purified to electrophoretic homogeneity. The sequences obtained from these clones could not be obtained directly either from the library or from total human DNA. The extended cDNA clones are not naturally occurring as such, but rather are obtained via manipulation of a partially purified naturally occurring substance (messenger RNA). The conversion of mRNA into a cDNA library involves the creation of a synthetic substance (cDNA) and pure individual cDNA clones can be isolated from the synthetic library by clonal selection. Thus, creating a cDNA library from messenger RNA and subsequently isolating individual clones from that library results in an approximately 10⁴-10⁶ fold purification of the native message. Purification of starting material or natural material to at least one order of magnitude, preferably two or three orders, and more preferably four or five orders of magnitude is expressly contemplated.

As used herein, the term "isolated" requires that the material be removed from its original environment (e.g., the natural environment if it is naturally occurring). For example, a naturally-occurring polynucleotide present in a living animal is not isolated, but the same polynucleotide, separated from some or all of the coexisting materials in the natural system, is isolated.

As used herein, the term "recombinant" means that the extended cDNA is adjacent to "backbone" nucleic acid to which it is not adjacent in its natural environment. Additionally, to be "enriched" the extended cDNAs will represent 5% or more of the number of nucleic acid inserts in a population of nucleic acid backbone molecules. Backbone molecules according to the present invention include nucleic acids such as expression vectors, self-replicating nucleic acids, viruses, integrating nucleic acids, and other vectors or nucleic acids used to maintain or manipulate a nucleic acid insert of interest. Preferably, the enriched extended cDNAs represent 15% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. More preferably, the enriched extended cDNAs represent 50% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. In a highly preferred embodiment, the enriched extended cDNAs represent 90% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. "Stringent", "moderate," and "low" hybridization conditions are as defined in Example 29.

Unless otherwise indicated, a "complementary" sequence is fully complementary. Thus, extended cDNAs encoding secreted polypeptides or fragments thereof which are present in cDNA libraries in which one or more extended cDNAs encoding secreted polypeptides or fragments thereof make up 5% or more of the number of nucleic acid inserts in the backbone molecules are "enriched recombinant extended cDNAs" as defined herein. Likewise, extended cDNAs encoding secreted polypeptides or fragments thereof which are in a population of plasmids in which one or more extended cDNAs of the present invention have been inserted such that they represent 5% or more of the number of inserts in the plasmid backbone are "enriched recombinant extended cDNAs" as defined herein. However, extended

cDNAs encoding secreted polypeptides or fragments thereof which are in cDNA libraries in which the extended cDNAs encoding secreted polypeptides or fragments thereof constitute less than 5% of the number of nucleic acid inserts in the population of backbone molecules, such as libraries in which backbone molecules having a cDNA insert encoding a secreted polypeptide are extremely rare, are not "enriched recombinant extended cDNAs."

In particular, the present invention relates to extended cDNAs which were derived from genes encoding secreted proteins. As used herein, a "secreted" protein is one which, when expressed in a suitable host cell, is transported across or through a membrane, including transport as a result of signal peptides in its amino acid sequence. "Secreted" proteins include without limitation proteins secreted wholly (e.g. soluble proteins), or partially (e.g. receptors) from the cell in which they are expressed. "Secreted" proteins also include without limitation proteins which are 10 transported across the membrane of the endoplasmic reticulum.

Extended cDNAs encoding secreted proteins may include nucleic acid sequences, called signal sequences, which encode signal puptides which direct the extracellular secretion of the proteins encoded by the extended cDNAs. Generally, the signal peptides are located at the amino termini of secreted proteins.

Secreted proteins are translated by ribosomes associated with the "rough" endoplasmic reticulum. Generally, 15 secreted proteins are co-translationally transferred to the membrane of the endoplasmic reticulum. Association of the ribosome with the endoplasmic reticulum during translation of secreted proteins is mediated by the signal peptide. The signal peptide is typically cleaved following its co-translational entry into the endoplasmic reticulum. After delivery to the endoplasmic reticulum, secreted proteins may proceed through the Golgi apparatus. In the Golgi apparatus, the proteins may undergo post-translational modification before entering secretory vesicles which transport them across the 20 cell membrane.

The extended cDNAs of the present invention have several important applications. For example, they may be used to express the entire secreted protein which they encode. Alternatively, they may be used to express portions of the secreted protein. The portions may comprise the signal peptides encoded by the extended cDNAs or the mature proteins encoded by the extended cDNAs (i.e. the proteins generated when the signal peptide is cleaved off). The 25 portions may also comprise polypeptides having at least 10 consecutive amino acids encoded by the extended cDNAs. Alternatively, the portions may comprise at least 15 consecutive amino acids encoded by the extended cDNAs. In some embodiments, the portions may comprise at least 25 consecutive amino acids encoded by the extended cDNAs. In other embodiments, the portions may comprise at least 40 amino acids encoded by the extended cDNAs.

Antibodies which specifically recognize the entire secreted proteins encoded by the extended cDNAs or 30 fragments thereof having at least 10 consecutive amino acids, at least 15 consecutive amino acids, at least 25 consecutive amino acids, or at least 40 consecutive amino acids may also be obtained as described below. Antibodies which specifically recognize the mature protein generated when the signal peptide is cleaved may also be obtained as described below. Similarly, antibodies which specifically recognize the signal peptides encoded by the extended cDNAs may also be obtained.

In some embodiments, the extended cDNAs include the signal sequence. In other embodiments, the extended cDNAs may include the full coding sequence for the mature protein (i.e. the protein generated when the signal polypeptide is cleaved off). In addition, the extended cDNAs may include regulatory regions upstream of the translation start site or downstream of the stop codon which control the amount, location, or developmental stage of gene expression. As discussed above, secreted proteins are therapeutically important. Thus, the proteins expressed from the cDNAs may be useful in treating or controlling a variety of human conditions. The extended cDNAs may also be used to obtain the corresponding genomic DNA. The term "corresponding genomic DNA" refers to the genomic DNA which encodes mRNA which includes the sequence of one of the strands of the extended cDNA in which thymidine residues in the sequence of the extended cDNA are replaced by uracil residues in the mRNA.

The extended cDNAs or genomic DNAs obtained therefrom may be used in forensic procedures to identify individuals or in diagnostic procedures to identify individuals having genetic diseases resulting from abnormal expression of the genes corresponding to the extended cDNAs. In addition, the present invention is useful for constructing a high resolution map of the human chromosomes.

The present invention also relates to secretion vectors capable of directing the secretion of a protein of

interest. Such vectors may be used in gene therapy strategies in which it is desired to produce a gene product in one cell
which is to be delivered to another location in the body. Secretion vectors may also facilitate the purification of desired
proteins.

The present invention also relates to expression vectors capable of directing the expression of an inserted gene in a desired spatial or temporal manner or at a desired level. Such vectors may include sequences upstream of the extended cDNAs such as promoters or upstream regulatory sequences.

In addition, the present invention may also be used for gene therapy to control or treat genetic diseases. Signal peptides may also be fused to heterologous proteins to direct their extracellular secretion.

One embodiment of the present invention is a purified or isolated nucleic acid comprising the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary thereto. In one aspect of this embodiment, the nucleic acid is recombinant.

Another embodiment of the present invention is a purified or isolated nucleic acid comprising at least 10 consecutive bases of the sequence of one of SEQ ID NOs: 40-140 and 242-377 or one of the sequences complementary thereto. In one aspect of this embodiment, the nucleic acid comprises at least 15, 25, 30, 40, 50, 75, or 100 consecutive bases of one of the sequences of SEQ ID NOs: 40-140 and 242-377 or one of the sequences complementary thereto. The nucleic acid may be a recombinant nucleic acid.

Another embodiment of the present invention is a purified or isolated nucleic acid of at least 15 bases capable of hybridizing under stringent conditions to the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary to one of the sequences of SEQ ID NOs: 40-140 and 242-377. In one aspect of this embodiment, the nucleic acid is recombinant.

25

30

Another embodiment of the present invention is a purified or isolated nucleic acid comprising the full coding sequences of one of SEQ ID NOs: 40-140 and 242-377, wherein the full coding sequence optionally comprises the sequence encoding signal peptide as well as the sequence encoding mature protein. In a preferred embodiment, the isolated or purified nucleic acid comprises the full coding sequence of one of SEQ ID Nos. 40, 42-44, 46, 48, 49, 51, 53, 60, 62-72, 76-78, 80-83, 85-88, 90, 93, 94, 97, 99-102, 104, 107-125, 127, 132, 135-138, 140 and 242-377 wherein the full coding sequence comprises the sequence encoding signal peptide and the sequence encoding mature protein. In one aspect of this embodiment, the nucleic acid is recombinant.

A further embodiment of the present invention is a purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode a mature protein. In a preferred embodiment, the purified or isolated nucleic acid comprises the nucleotides of one of SEQ ID NOs: 40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode a mature protein. In one aspect of this embodiment, the nucleic acid is recombinant.

Yet another embodiment of the present invention is a purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode the signal peptide. In a preferred embodiment, the purified or isolated nucleic acid comprises the nucleotides of SEQ ID NOs: 40, 42-46, 48, 49, 51, 53, 57, 60, 62-73, 76-78, 80-83, 85-88, 90, 93-95, 97, 99-102, 104, 107-125, 127, 128, 130, 132, 134-140 and 242-377 which encode the signal peptide. In one aspect of this embodiment, the nucleic acid is recombinant.

Another embodiment of the present invention is a purified or isolated nucleic acid encoding a polypeptide having the sequence of one of the sequences of SEQ ID NOs: 141-241 and 378-513.

Another embodiment of the present invention is a purified or isolated nucleic acid encoding a polypeptide having the sequence of a mature protein included in one of the sequences of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the purified or isolated nucleic acid encodes a polypeptide having the sequence of a mature protein included in one of the sequences of SEQ ID NOs: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.

Another embodiment of the present invention is a purified or isolated nucleic acid encoding a polypeptide having the sequence of a signal peptide included in one of the sequences of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the purified or isolated nucleic acid encodes a polypeptide having the sequence of a signal peptide included in one of the sequences of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-226, 228, 229, 231, 233, 235-241, and 378-513.

Yet another embodiment of the present invention is a purified or isolated protein comprising the sequence of one of SEQ ID NOs: 141-241 and 378-513.

Another embodiment of the present invention is a purified or isolated polypeptide comprising at least 10 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In one aspect of this embodiment, the purified or isolated polypeptide comprises at least 15, 20, 25, 35, 50, 75, 100, 150 or 200 consecutive

amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In still another aspect, the purified or isolated polypeptide comprises at least 25 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513.

Another embodiment of the present invention is an isolated or purified polypeptide comprising a signal peptide of one of the polypeptides of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the isolated or purified polypeptide comprises a signal peptide of one of the polypeptides of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-226, 228, 229, 231, 233, 235-241, and 378-513.

Yet another embodiment of the present invention is an isolated or purified polypeptide comprising a mature protein of one of the polypeptides of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the isolated or purified polypeptide comprises a mature protein of one of the polypeptides of SEQ ID NOs: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.

A further embodiment of the present invention is a method of making a protein comprising one of the sequences of SEQ ID NO: 141-241 and 378-513, comprising the steps of obtaining a cDNA comprising one of the sequences of sequence of SEQ ID NO: 40-140 and 242-377, inserting the cDNA in an expression vector such that the cDNA is operably linked to a promoter, and introducing the expression vector into a host cell whereby the host cell produces the protein encoded by said cDNA. In one aspect of this embodiment, the method further comprises the step of isolating the protein.

Another embodiment of the present invention is a protein obtainable by the method described in the preceding paragraph.

Another embodiment of the present invention is a method of making a protein comprising the amino acid sequence of the mature protein contained in one of the sequences of SEQ ID NO: 141-241 and 378-513, comprising the steps of obtaining a cDNA comprising one of the nucleotides sequence of sequence of SEQ ID NO: 40-140 and 242-377 which encode for the mature protein, inserting the cDNA in an expression vector such that the cDNA is operably linked to a promoter, and introducing the expression vector into a host cell whereby the host cell produces the mature protein encoded by the cDNA. In one aspect of this embodiment, the method further comprises the step of isolating the protein.

Another embodiment of the present invention is a mature protein obtainable by the method described in the preceding paragraph.

In a preferred embodiment, the above method comprises a method of making a protein comprising the amino acid sequence of the mature protein contained in one of the sequences of SEQ ID NO: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513, comprising the steps of obtaining a cDNA comprising one of the nucleotides sequence of sequence of SEQ ID NO:

40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode for the mature protein, inserting the cDNA in an expression vector such that the cDNA is operably linked to a promoter, and introducing the expression vector into a host cell whereby the host cell produces the mature protein encoded by the cDNA. In one aspect of this embodiment, the method further comprises the step of isolating the protein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary thereto described herein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the full coding sequences of one of SEQ ID NOs: 40-140 and 242-377, wherein the full coding sequence comprises the sequence encoding signal peptide and the sequence encoding mature protein described herein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode a mature protein which are described herein. Preferably, the host cell contains the purified or isolated nucleic acids comprising the nucleotides of one of SEQ ID NOs: 40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode a mature protein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode the signal peptide which are described herein. Preferably, the host cell contains the purified or isolated nucleic acids comprising the nucleotides of one of SEQ ID Nos.: 40, 42-46, 48, 49, 51, 53, 57, 60, 62-73, 76-78, 80-83, 85-88, 90, 93-95, 97, 99-102, 104, 107-125, 127, 128, 130, 132, 134-140 and 242-377 which encode the signal peptide.

Another embodiment of the present invention is a purified or isolated antibody capable of specifically binding to a protein having the sequence of one of SEQ ID NOs: 141-241 and 378-513. In one aspect of this embodiment, the antibody is capable of binding to a polypeptide comprising at least 10 consecutive amino acids of the sequence of one of SEQ ID NOs: 141-241 and 378-513.

Another embodiment of the present invention is an array of cDNAs or fragments thereof of at least 15 nucleotides in length which includes at least one of the sequences of SEQ ID NOs: 40-140 and 242-377, or one of the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or a fragment thereof of at least 15 consecutive nucleotides. In one aspect of this embodiment, the array includes at least two of the sequences of SEQ ID NOs: 40-140 and 242-377, the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or fragments thereof of at least 15 consecutive nucleotides. In another aspect of this embodiment, the array includes at least five of the sequences of SEQ ID NOs: 40-140 and 242-377, the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or fragments thereof of at least 15 consecutive nucleotides.

A further embodiment of the invention encompasses purified polynucleotides comprising an insert from a clone deposited in a deposit having an accession number selected from the group consisting of the accession numbers listed in Table VI or a fragment thereof comprising a contiguous span of at least 8, 10, 12, 15, 20, 25, 40, 60, 100, or 200 nucleotides of said insert. An additional embodiment of the invention encompasses purified polypeptides which comprise, consist of, or consist essentially of an amino acid sequence encoded by the insert from a clone deposited in a deposit having an accession number selected from the group consisting of the accession numbers listed in Table VI, as well as polypeptides which comprise a fragment of said amino acid sequence consisting of a signal peptide, a mature protein, or a contiguous span of at least 5, 8, 10, 12, 15, 20, 25, 40, 60, 100, or 200 amino acids encoded by said insert.

An additional embodiment of the invention encompasses purified polypeptides which comprise a contiguous span of at least 5, 8, 10, 12, 15, 20, 25, 40, 60, 100, or 200 amino acids of SEQ ID NOs: 158, 174, 175, 196, 226, 231, 232, wherein said contiguous span comprises at least one of the amino acid positions which was not shown to be identical to a public sequence in any of Figures 11 to 15. Also encompassed by the invention are purified polynoculeotides encoding said polypeptides.

15

10

Brief Description of the Drawings

Figure 1 is a summary of a procedure for obtaining cDNAs which have been selected to include the 5' ends of the mRNAs from which they are derived.

Figure 2 is an analysis of the 43 amino terminal amino acids of all human SwissProt proteins to determine the frequency of false positives and false negatives using the techniques for signal peptide identification described herein.

Figure 3 shows the distribution of von Heijne scores for 5' ESTs in each of the categories described herein and the probability that these 5' ESTs encode a signal peptide.

Figure 4 shows the distribution of 5' ESTs in each category and the number of 5' ESTs in each category having a given minimum von Heijne's score.

Figure 5 shows the tissues from which the mRNAs corresponding to the 5' ESTs in each of the categories described herein were obtained.

Figure 6 illustrates a method for obtaining extended cDNAs.

Figure 7 is a map of pED6dpc2. pED6dpc2 is derived from pED6dpc1 by insertion of a new polylinker to facilitate cDNA cloning. SSt cDNAs are cloned between EcoRI and NotI. PED vectors are described in Kaufman et al. 30 (1991), NAR 19: 4485-4490.

Figure 8 provides a schematic description of the promoters isolated and the way they are assembled with the corresponding 5' tags.

Figure 9 describes the transcription factor binding sites present in each of these promoters.

Figure 10 is an alignment of the protein of SEQ ID NO: 217 with the human protein TFAR19 that may play a role in apoptosis (Genbank accession number AF014955, SEQ ID NO: 516).

Figure 11 is an alignment of the proteins of SEQ ID NOs: 174, 175 and 232 with a human secreted protein (Genseg accession number W36955, SEQ ID NO: 517).

Figure 12 is an alignment of the protein of SEQ ID NO: 231 with the human E25 protein (Genbank accession number AF038953, SEQ ID NO: 515).

Figure 13 is an alignment of the protein of SEQ ID NO: 196 with the human seventransmembrane protein (Genbank accession number Y11395, SEQ ID NO: 518).

Figure 14 is an alignment of the protein of SEQ ID NOs: 158 with the murine subunit 7a of the COP9 complex 10 (Genbank accession number AF071316, SEQ ID NO: 519).

Figure 15 is an alignment of the protein of SEQ ID NO: 226 with the bovine subunit B14.5B of the NADHubiquinone oxidureductase complex (Arizmendi *et al, FEBS Lett.*, 313: 80-84 (1992) and Swissprot accession -number Q02827, SEQ ID NO: 514).

Detailed Description of the Preferred Embodiment

15 I. Obtaining 5' ESTs

5

The present extended cDNAs were obtained using 5' ESTs which were isolated as described below.

A. Chemical Methods for Obtaining mRNAs having Intact 5' Ends

In order to obtain the 5' ESTs used to obtain the extended cDNAs of the present invention, mRNAs having intact 5' ends must be obtained. Currently, there are two approaches for obtaining such mRNAs. One of these 20 approaches is a chemical modification method involving derivatization of the 5' ends of the mRNAs and selection of the derivatized mRNAs. The 5' ends of eucaryotic mRNAs possess a structure referred to as a "cap" which comprises a guanosine methylated at the 7 position. The cap is joined to the first transcribed base of the mRNA by a 5', 5'triphosphate bond. In some instances, the 5' guanosine is methylated in both the 2 and 7 positions. Rarely, the 5' guanosine is trimethylated at the 2, 7 and 7 positions. In the chemical method for obtaining mRNAs having intact 5' 25 ends, the 5' cap is specifically derivatized and coupled to a reactive group on an immobilizing substrate. This specific derivatization is based on the fact that only the ribose linked to the methylated guanosine at the 5' end of the mRNA and the ribose linked to the base at the 3' terminus of the mRNA, possess 2', 3'-cis diols. Optionally, where the 3' terminal ribose has a 2', 3'-cis diol, the 2', 3'-cis diol at the 3' end may be chemically modified, substituted, converted, or eliminated, leaving only the ribose linked to the methylated guanosine at the 5' end of the mRNA with a 2', 3'-cis diol. A 30 variety of techniques are available for eliminating the 2', 3'-cis diol on the 3' terminal ribose. For example, controlled alkaline hydrolysis may be used to generate mRNA fragments in which the 3' terminal ribose is a 3'-phosphate, 2'phosphate or (2', 3')-cyclophosphate. Thereafter, the fragment which includes the original 3' ribose may be eliminated from the mixture through chromatography on an oligo-dT column. Alternatively, a base which lacks the 2', 3'-cis diol

may be added to the 3' end of the mRNA using an RNA ligase such as T4 RNA ligase. Example 1 below describes a method for ligation of pCp to the 3' end of messenger RNA.

EXAMPLE 1

Ligation of the Nucleoside Diphosphate pCp to the 3' End of Messenger RNA

 $1 \mu g$ of RNA was incubated in a final reaction medium of 10 μl in the presence of 5 U of T₄ phage RNA ligase in the buffer provided by the manufacturer (Gibco - BRL), 40 U of the RNase inhibitor RNasin (Promega) and, 2 μl of 32 pCp (Amersham #PB 10208).

The incubation was performed at 37°C for 2 hours or overnight at 7-8°C.

Following modification or elimination of the 2', 3'-cis diol at the 3' ribose, the 2', 3'-cis diol present at the 5' end of the mRNA may be oxidized using reagents such as NaBH₄, NaBH₃CN, or sodium periodate, thereby converting the 2', 3'-cis diol to a dialdehyde. Example 2 describes the oxidation of the 2', 3'-cis diol at the 5' end of the mRNA with sodium periodate.

EXAMPLE 2

Oxidation of 2', 3'-cis diol at the 5' End of the mRNA

O.1 OD unit of either a capped oligoribonucleotide of 47 nucleotides (including the cap) or an uncapped oligoribonucleotide of 46 nucleotides were treated as follows. The oligoribonucleotides were produced by in vitro transcription using the transcription kit "AmpliScribe T7" (Epicentre Technologies). As indicated below, the DNA template for the RNA transcript contained a single cytosine. To synthesize the uncapped RNA, all four NTPs were included in the in vitro transcription reaction. To obtain the capped RNA, GTP was replaced by an analogue of the cap, m7G(5')ppp(5')G. This compound, recognized by polymerase, was incorporated into the 5' end of the nascent transcript during the step of initiation of transcription but was not capable of incorporation during the extension step.

Consequently, the resulting RNA contained a cap at its 5' end. The sequences of the oligoribonucleotides produced by the in vitro transcription reaction were:

+ Cap:

25 5'm7GpppGCAUCCUACUCCCAUCCAAUUCCACCCUAACUCCUCCCAUCUCCAC-3' (SEQ ID NO:1)

-Cap:

5'-pppGCAUCCUACUCCCAUCCAAUUCCACCCUAACUCCUCCCAUCUCCAC-3' (SEQ ID NO:2)

The oligoribonucleotides were dissolved in 9 µl of acetate buffer (0.1 M sodium acetate, pH 5.2) and 3 µl of freshly prepared 0.1 M sodium periodate solution. The mixture was incubated for 1 hour in the dark at 4°C or room temperature. Thereafter, the reaction was stopped by adding 4 µl of 10% ethylene glycol. The product was ethanol precipitated, resuspended in 10µl or more of water or appropriate buffer and dialyzed against water.

The resulting aldehyde groups may then be coupled to molecules having a reactive amine group, such as hydrazine, carbazide, thiocarbazide or semicarbazide groups, in order to facilitate enrichment of the 5' ends of the mRNAs. Molecules having reactive amine groups which are suitable for use in selecting mRNAs having intact 5' ends

include avidin, proteins, antibodies, vitamins, ligands capable of specifically binding to receptor molecules, or oligonucleotides. Example 3 below describes the coupling of the resulting dialdehyde to biotin.

EXAMPLE 3

Coupling of the Dialdehyde with Biotin

The oxidation product obtained in Example 2 was dissolved in 50 μl of sodium acetate at a pH of between 5 and 5.2 and 50 μl of freshly prepared 0.02 M solution of biotin hydrazide in a methoxyethanol/water mixture (1:1) of formula:

In the compound used in these experiments, n=5. However, it will be appreciated that other commercially available hydrazides may also be used, such as molecules of the formula above in which n varies from 0 to 5.

The mixture was then incubated for 2 hours at 37°C. Following the incubation, the mixture was precipitated with ethanol and dialyzed against distilled water.

Example 4 demonstrates the specificity of the biotinylation reaction.

15

EXAMPLE 4

Specificity of Biotinylation

The specificity of the biotinylation for capped mRNAs was evaluated by gel electrophoresis of the following samples:

- Sample 1. The 46 nucleotide uncapped in vitro transcript prepared as in Example 2 and labeled with ³²pCp as 20 described in Example 1.
 - Sample 2. The 46 nucleotide uncapped in vitro transcript prepared as in Example 2, labeled with ³²pCp as described in Example 1, treated with the oxidation reaction of Example 2, and subjected to the biotinylation conditions of Example 3.
- Sample 3. The 47 nucleotide capped in vitro transcript prepared as in Example 2 and labeled with ³²pCp as described in Example 1.
 - Sample 4. The 47 nucleotide capped in vitro transcript prepared as in Example 2, labeled with ³²pCp as described in Example 1, treated with the oxidation reaction of Example 2, and subjected to the biotinylation conditions of Example 3.
- Samples 1 and 2 had indentical migration rates, demonstrating that the uncapped RNAs were not oxidized and biotinylated. Sample 3 migrated more slowly than Samples 1 and 2, while Sample 4 exhibited the slowest migration.

25

The difference in migration of the RNAs in Samples 3 and 4 demonstrates that the capped RNAs were specifically biotinylated.

In some cases, mRNAs having intact 5' ends may be enriched by binding the molecule containing a reactive amine group to a suitable solid phase substrate such as the inside of the vessel containing the mRNAs, magnetic beads, 5 chromatography matrices, or nylon or nitrocellulose membranes. For example, where the molecule having a reactive amine group is biotin, the solid phase substrate may be coupled to avidin or streptavidin. Alternatively, where the molecule having the reactive amine group is an antibody or receptor ligand, the solid phase substrate may be coupled to the cognate antigen or receptor. Finally, where the molecule having a reactive amine group comprises an oligonucleotide, the solid phase substrate may comprise a complementary oligonucleotide.

The mRNAs having intact 5' ends may be released from the solid phase following the enrichment procedure. For example, where the dialdehyde is coupled to biotin hydrazide and the solid phase comprises streptavidin, the mRNAs may be released from the solid phase by simply heating to 95 degrees Celsius in 2% SDS. In some methods, the molecule having a reactive amine group may also be cleaved from the mRNAs having intact 5' ends following enrichment. Example 5 describes the capture of biotinylated mRNAs with streptavidin coated beads and the release of the 15 biotinylated mRNAs from the beads following enrichment.

EXAMPLE 5

Capture and Release of Biotinylated mRNAs Using Strepatividin Coated Beads

The streptavidin-coated magnetic beads were prepared according to the manufacturer's instructions (CPG Inc., USA). The biotinylated mRNAs were added to a hybridization buffer (1.5 M NaCl, pH 5 · 6). After incubating for 30 20 minutes, the unbound and nonbiotinylated material was removed. The beads were washed several times in water with 1% SDS. The beads obtained were incubated for 15 minutes at 95°C in water containing 2% SDS.

Example 6 demonstrates the efficiency with which biotinylated mRNAs were recovered from the streptavidin coated beads.

EXAMPLE 6

Efficiency of Recovery of Biotinylated mRNAs

The efficiency of the recovery procedure was evaluated as follows. RNAs were labeled with ³²pCp, oxidized, biotinylated and bound to streptavidin coated beads as described above. Subsequently, the bound RNAs were incubated for 5, 15 or 30 minutes at 95°C in the presence of 2% SDS.

The products of the reaction were analyzed by electrophoresis on 12% polyacrylamide gels under denaturing 30 conditions (7 M urea). The gels were subjected to autoradiography. During this manipulation, the hydrazone bonds were not reduced.

Increasing amounts of nucleic acids were recovered as incubation times in 2% SDS increased, demonstrating that biotinylated mRNAs were efficiently recovered.

In an alternative method for obtaining mRNAs having intact 5' ends, an oligonucleotide which has been derivatized to contain a reactive amine group is specifically coupled to mRNAs having an intact cap. Preferably, the 3' end of the mRNA is blocked prior to the step in which the aldehyde groups are joined to the derivatized oligonucleotide, as described above, so as to prevent the derivatized oligonucleotide from being joined to the 3' end of the mRNA. For example, pCp may be attached to the 3' end of the mRNA using T4 RNA ligase. However, as discussed above, blocking the 3' end of the mRNA is an optional step. Derivatized oligonucleotides may be prepared as described below in Example 7.

EXAMPLE 7

Derivatization of the Oligonucleotide

An oligonucleotide phosphorylated at its 3' end was converted to a 3' hydrazide in 3' by treatment with an aqueous solution of hydrazine or of dihydrazide of the formula $H_2N(R1)NH_2$ at about 1 to 3 M, and at pH 4.5, in the presence of a carbodiimide type agent soluble in water such as 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide at a final concentration of 0.3 M at a temperature of 8°C overnight.

The derivatized oligonucleotide was then separated from the other agents and products using a standard technique for isolating oligonucleotides.

As discussed above, the mRNAs to be enriched may be treated to eliminate the 3' OH groups which may be present thereon. This may be accomplished by enzymatic ligation of sequences lacking a 3' OH, such as pCp, as described above in Example 1. Alternatively, the 3' OH groups may be eliminated by alkaline hydrolysis as described in Example 8 below.

20

25

EXAMPLE 8

Alkaline Hydrolysis of mRNA

The mRNAs may be treated with alkaline hydrolysis as follows. In a total volume of 100µl of 0.1N sodium hydroxide, 1.5µg mRNA is incubated for 40 to 60 minutes at 4°C. The solution is neutralized with acetic acid and precipitated with ethanol.

Following the optional elimination of the 3' OH groups, the diol groups at the 5' ends of the mRNAs are oxidized as described below in Example 9.

EXAMPLE 9

Oxidation of Diols

Up to 1 OD unit of RNA was dissolved in 9 µl of buffer (0.1 M sodium acetate, pH 6-7 or water) and 3 µl of freshly prepared 0.1 M sodium periodate solution. The reaction was incubated for 1 h in the dark at 4°C or room temperature. Following the incubation, the reaction was stopped by adding 4 µl of 10% ethylene glycol. Thereafter the mixture was incubated at room temperature for 15 minutes. After ethanol precipitation, the product was resuspended in 10µl or more of water or appropriate buffer and dialyzed against water.

Following oxidation of the diol groups at the 5' ends of the mRNAs, the derivatized oligonucleotide was joined to the resulting aldehydes as described in Example 10.

EXAMPLE 10

Reaction of Aldehydes with Derivatized Oligonucleotides

The oxidized mRNA was dissolved in an acidic medium such as 50 µl of sodium acetate pH 4-6. 50 µl of a solution of the derivatized oligonucleotide was added such that an mRNA:derivatized oligonucleotide ratio of 1:20 was obtained and mixture was reduced with a borohydride. The mixture was allowed to incubate for 2 h at 37°C or overnight (14 h) at 10°C. The mixture was ethanol precipitated, resuspended in 10µl or more of water or appropriate buffer and dialyzed against distilled water. If desired, the resulting product may be analyzed using acrylamide gel electrophoresis, HPLC analysis, or other conventional techniques.

Following the attachment of the derivatized oligonucleotide to the mRNAs, a reverse transcription reaction may be performed as described in Example 11 below.

EXAMPLE 11

Reverse Transcription of mRNAs

An oligodeoxyribonucleotide was derivatized as follows. 3 OD units of an oligodeoxyribonucleotide of sequence ATCAAGAATTCGCACGAGACCATTA (SEQ ID NO:3) having 5'-OH and 3'-P ends were dissolved in 70 µl of a 1.5 M hydroxybenzotriazole solution, pH 5.3, prepared in dimethylformamide/water (75:25) containing 2 µg of 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide. The mixture was incubated for 2 h 30 min at 22°C. The mixture was then precipitated twice in LiClO₄/acetone. The pellet was resuspended in 200 µl of 0.25 M hydrazine and incubated at 8°C from 3 to 14 h. Following the hydrazine reaction, the mixture was precipitated twice in LiClO₄/acetone.

The messenger RNAs to be reverse transcribed were extracted from blocks of placenta having sides of 2 cm which had been stored at -80°C. The mRNA was extracted using conventional acidic phenol techniques. Oligo-dT chromatography was used to purify the mRNAs. The integrity of the mRNAs was checked by Northern-blotting.

The diol groups on 7 µg of the placental mRNAs were oxidized as described above in Example 9. The

derivatized oligonucleotide was joined to the mRNAs as described in Example 10 above except that the precipitation step was replaced by an exclusion chromatography step to remove derivatized oligodeoxyribonucleotides which were not joined to mRNAs. Exclusion chromatography was performed as follows:

10 ml of AcA34 (BioSepra#230151) gel were equilibrated in 50 ml of a solution of 10 mM Tris pH 8.0, 300 mM NaCl, 1 mM EDTA, and 0.05% SDS. The mixture was allowed to sediment. The supernatant was eliminated and the gel was resuspended in 50 ml of buffer. This procedure was repeated 2 or 3 times.

A glass bead (diameter 3 mm) was introduced into a 2 ml disposable pipette (length 25 cm). The pipette was filled with the gel suspension until the height of the gel stabilized at 1 cm from the top of the pipette. The column was then equilibrated with 20 ml of equilibration buffer (10 mM Tris HCl pH 7.4, 20 mM NaCl).

. .. .

10 μ l of the mRNA which had been reacted with the derivatized oligonucleotide were mixed in 39 μ l of 10 mM urea and 2 μ l of blue-glycerol buffer, which had been prepared by dissolving 5 mg of bromophenol blue in 60% glycerol (v/v), and passing the mixture through a filter with a filter of diameter 0.45 μ m.

The column was loaded. As soon as the sample had penetrated, equilibration buffer was added. 100 µl fractions were collected. Derivatized oligonucleotide which had not been attached to mRNA appeared in fraction 16 and later fractions. Fractions 3 to 15 were combined and precipitated with ethanol.

The mRNAs which had been reacted with the derivatized oligonucleotide were spotted on a nylon membrane and hybridized to a radioactive probe using conventional techniques. The radioactive probe used in these hybridizations was an oligodeoxyribonucleotide of sequence TAATGGTCTCGTGCGAATTCTTGAT (SEQ ID NO:4) which was anticomplementary to the derivatized oligonucleotide and was labeled at its 5' end with ³²P. 1/10th of the mRNAs which had been reacted with the derivatized oligonucleotide was spotted in two spots on the membrane and the membrane was visualized by autoradiography after hybridization of the probe. A signal was observed, indicating that the derivatized oligonucleotide had been joined to the mRNA.

The remaining 9/10 of the mRNAs which had been reacted with the derivatized oligonucleotide was reverse transcribed as follows. A reverse transcription reaction was carried out with reverse transcriptase following the manufacturer's instructions. To prime the reaction, 50 pmol of nonamers with random sequence were used.

A portion of the resulting cDNA was spotted on a positively charged nylon membrane using conventional methods. The cDNAs were spotted on the membrane after the cDNA:RNA heteroduplexes had been subjected to an alkaline hydrolysis in order to eliminate the RNAs. An oligonucleotide having a sequence identical to that of the derivatized oligonucleotide was labeled at its 5' end with ³²P and hybridized to the cDNA blots using conventional techniques. Single-stranded cDNAs resulting from the reverse transcription reaction were spotted on the membrane. As controls, the blot contained 1 pmol, 100 fmol, 50 fmol, 10 fmol and 1 fmol respectively of a control oligodeoxyribonucleotide of sequence identical to that of the derivatized oligonucleotide. The signal observed in the spots containing the cDNA indicated that approximately 15 fmol of the derivatized oligonucleotide had been reverse transcribed.

These results demonstrate that the reverse transcription can be performed through the cap and, in particular, that reverse transcriptase crosses the 5'-P-P-5' bond of the cap of eukaryotic messenger RNAs.

The single stranded cDNAs obtained after the above first strand synthesis were used as template for PCR reactions. Two types of reactions were carried out. First, specific amplification of the mRNAs for the alpha globin, dehydrogenase, pp15 and elongation factor E4 were carried out using the following pairs of oligodeoxyribonucleotide primers.

alpha-globin

25

GLO-S: CCG ACA AGA CCA ACG TCA AGG CCG C (SEQ ID NO:5)

GLO-As: TCA CCA GCA GGC AGT GGC TTA GGA G 3' (SEQ ID NO:6)

dehydrogenase

3 DH-S: AGT GAT TCC TGC TAC TTT GGA TGG C (SEQ ID NO:7)

3 DH-As: GCT TGG TCT TGT TCT GGA GTT TAG A (SEQ ID NO:8)

15מם

PP15-S: TCC AGA ATG GGA GAC AAG CCA ATT T (SEQ ID NO:9)

5 PP15-As: AGG GAG GAG GAA ACA GCG TGA GTC C (SEQ ID NO:10)

Elongation factor E4

EFA1-S: ATG GGA AAG GAA AAG ACT CAT ATC A (SEQ ID NO:11)

EF1A-As: AGC AGC AAC AAT CAG GAC AGC ACA G (SEQ ID NO:12)

Non specific amplifications were also carried out with the antisense (_As) oligodeoxyribonucleotides of the pairs described above and a primer chosen from the sequence of the derivatized oligodeoxyribonucleotide (ATCAAGAATTCGCACGAGACCATTA) (SEQ ID NO:13).

A 1.5% agarose gel containing the following samples corresponding to the PCR products of reverse transcription was stained with ethidium bromide. (1/20th of the products of reverse transcription were used for each PCR reaction).

- Sample 1: The products of a PCR reaction using the globin primers of SEQ ID NOs 5 and 6 in the presence of cDNA.
 - Sample 2: The products of a PCR reaction using the globin primers of SEQ ID NOs 5 and 6 in the absence of added cDNA.
- Sample 3: The products of a PCR reaction using the dehydrogenase primers of SEQ ID NOs 7 and 8 in the presence of cDNA.
 - Sample 4: The products of a PCR reaction using the dehydrogenase primers of SEO ID NOs 7 and 8 in the absence of added cDNA.
 - Sample 5: The products of a PCR reaction using the pp15 primers of SEQ ID NOs 9 and 10 in the presence of cDNA.
- 25 Sample 6: The products of a PCR reaction using the pp15 primers of SEQ ID NOs 9 and 10 in the absence of added cDNA.
 - Sample 7: The products of a PCR reaction using the EIE4 primers of SEQ ID NOs 11 and 12 in the presence of added cDNA.
- Sample 8: The products of a PCR reaction using the EIE4 primers of SEQ ID NOs 11 and 12 in the absence of 30 added cDNA.

In Samples 1, 3, 5 and 7, a band of the size expected for the PCR product was observed, indicating the presence of the corresponding sequence in the cDNA population.

PCR reactions were also carried out with the antisense oligonucleotides of the globin and dehydrogenase primers (SEQ ID NOs 6 and 8) and an oligonucleotide whose sequence corresponds to that of the derivatized

oligonucleotide. The presence of PCR products of the expected size in the samples corresponding to samples 1 and 3 above indicated that the derivatized oligonucleotide had been incorporated.

The above examples summarize the chemical procedure for enriching mRNAs for those having intact 5' ends.

Further detail regarding the chemical approaches for obtaining mRNAs having intact 5' ends are disclosed in

International Application No. W096/34981, published November 7, 1996.

Strategies based on the above chemical modifications to the 5' cap structure may be utilized to generate cDNAs which have been selected to include the 5' ends of the mRNAs from which they are derived. In one version of such procedures, the 5' ends of the mRNAs are modified as described above. Thereafter, a reverse transcription reaction is conducted to extend a primer complementary to the mRNA to the 5' end of the mRNA. Single stranded RNAs are eliminated to obtain a population of cDNA/mRNA heteroduplexes in which the mRNA includes an intact 5' end. The resulting heteroduplexes may be captured on a solid phase coated with a molecule capable of interacting with the molecule used to derivatize the 5' end of the mRNA. Thereafter, the strands of the heteroduplexes are separated to recover single stranded first cDNA strands which include the 5' end of the mRNA. Second strand cDNA synthesis may then proceed using conventional techniques. For example, the procedures disclosed in WO 96/34981 or in Carninci, P. et al. High-Efficiency Full-Length cDNA Cloning by Biotinylated CAP Trapper. Genomics 37:327-336 (1996) may be employed to select cDNAs which include the sequence derived from the 5' end of the coding sequence of the mRNA.

Following ligation of the oligonucleotide tag to the 5' cap of the mRNA, a reverse transcription reaction is conducted to extend a primer complementary to the mRNA to the 5' end of the mRNA. Following elimination of the RNA component of the resulting heteroduplex using standard techniques, second strand cDNA synthesis is conducted with a 20 primer complementary to the oligonucleotide tag.

Figure 1 summarizes the above procedures for obtaining cDNAs which have been selected to include the 5' ends of the mRNAs from which they are derived.

B. Enzymatic Methods for Obtaining mRNAs having Intact 5' Ends

Other techniques for selecting cDNAs extending to the 5' end of the mRNA from which they are derived are
fully enzymatic. Some versions of these techniques are disclosed in Dumas Milne Edwards J.B. (Doctoral Thesis of Paris
VI University, Le clonage des ADNc complets: difficultes et perspectives nouvelles. Apports pour l'etude de la regulation
de l'expression de la tryptophane hydroxylase de rat, 20 Dec. 1993), EPO 625572 and Kato et al. Construction of a
Human Full-Length cDNA Bank. Gene 150:243-250 (1994).

Briefly, in such approaches, isolated mRNA is treated with alkaline phosphatase to remove the phosphate

30 groups present on the 5' ends of uncapped incomplete mRNAs. Following this procedure, the cap present on full length mRNAs is enzymatically removed with a decapping enzyme such as T4 polynucleotide kinase or tobacco acid pyrophosphatase. An oligonucleotide, which may be either a DNA oligonucleotide or a DNA-RNA hybrid oligonucleotide having RNA at its 3' end, is then ligated to the phosphate present at the 5' end of the decapped mRNA using T4 RNA

ligase. The oligonucleotide may include a restriction site to facilitate cloning of the cDNAs following their synthesis. Example 12 below describes one enzymatic method based on the doctoral thesis of Dumas.

EXAMPLE 12

Enzymatic Approach for Obtaining 5' ESTs

Twenty micrograms of PolyA+ RNA were dephosphorylated using Calf Intestinal Phosphatase (Biolabs). After a phenol chloroform extraction, the cap structure of mRNA was hydrolysed using the Tobacco Acid Pyrophosphatase (purified as described by Shinshi et al., Biochemistry 15: 2185-2190, 1976) and a hemi 5'DNA/RNA-3' oligonucleotide having an unphosphorylated 5' end, a stretch of adenosine ribophosphate at the 3' end, and an EcoRI site near the 5' end was ligated to the 5'P ends of mRNA using the T4 RNA ligase (Biolabs). Oligonucleotides suitable for use in this 10 procedure are preferably 30-50 bases in length. Oligonucleotides having an unphosphorylated 5' end may be synthesized by adding a fluorochrome at the 5' end. The inclusion of a stretch of adenosine ribophosphates at the 3' end of the oligonucleotide increases ligation efficiency. It will be appreciated that the oligonucleotide may contain cloning sites other than EcoRI.

Following ligation of the oligonucleotide to the phosphate present at the 5' end of the decapped mRNA, first 15 and second strand cDNA synthesis may be carried out using conventional methods or those specified in EPO 625,572 and Kato et al. Construction of a Human Full-Length cDNA Bank. Gene 150:243-250 (1994), and Dumas Milne Edwards, supra. The resulting cDNA may then be ligated into vectors such as those disclosed in Kato et al. Construction of a Human Full-Length cDNA Bank. Gene 150:243-250 (1994) or other nucleic acid vectors known to those skilled in the art using techniques such as those described in Sambrook et al., Molecular Cloning: A Laboratory Manual 2d Ed., Cold 20 Spring Harbor Laboratory Press, 1989.

II. Characterization of 5' ESTs

5

The above chemical and enzymatic approaches for enriching mRNAs having intact 5' ends were employed to obtain 5' ESTs. First, mRNAs were prepared as described in Example 13 below.

EXAMPLE 13

25 Preparation of mRNA

Total human RNAs or PolyA + RNAs derived from 29 different tissues were respectively purchased from LABIMO and CLONTECH and used to generate 44 cDNA libraries as described below. The purchased RNA had been isolated from cells or tissues using acid quanidium thiocyanate-phenol-chloroform extraction (Chomczyniski, P and Sacchi, N., Analytical Biochemistry 162:156-159, 1987). PolyA+ RNA was isolated from total RNA (LABIMO) by 30 two passes of oligodT chromatography, as described by Aviv and Leder (Aviv, H. and Leder, P., Proc. Natl. Acad. Sci. **USA** 69:1408-1412, 1972) in order to eliminate ribosomal RNA.

The quality and the integrity of the poly A+ were checked. Northern blots hybridized with a globin probe were used to confirm that the mRNAs were not degraded. Contamination of the PolyA + mRNAs by ribosomal sequences was checked using RNAs blots and a probe derived from the sequence of the 28S RNA. Preparations of mRNAs with less

than 5% of ribosomal RNAs were used in library construction. To avoid constructing libraries with RNAs contaminated by exogenous sequences (prokaryotic or fungal), the presence of bacterial 16S ribosomal sequences or of two highly expressed mRNAs was examined using PCR.

Following preparation of the mRNAs, the above described chemical and/or the enzymatic procedures for enriching mRNAs having intact 5' ends discussed above were employed to obtain 5' ESTs from various tissues. In both approaches an oligonucleotide tag was attached to the cap at the 5' ends of the mRNAs. The oligonucleotide tag had an EcoRI site therein to facilitate later cloning procedures.

Following attachment of the oligonucleotide tag to the mRNA by either the chemical or enzymatic methods, the integrity of the mRNA was examined by performing a Northern blot with 200-500ng of mRNA using a probe

10 complementary to the oligonucleotide tag.

EXAMPLE 14

cDNA Synthesis Using mRNA Templates Having Intact 5' Ends

For the mRNAs joined to oligonucleotide tags using both the chemical and enzymatic methods, first strand cDNA synthesis was performed using reverse transcriptase with random nonamers as primers. In order to protect internal EcoRI sites in the cDNA from digestion at later steps in the procedure, methylated dCTP was used for first strand synthesis. After removal of RNA by an alkaline hydrolysis, the first strand of cDNA was precipitated using isopropanol in order to eliminate residual primers.

For both the chemical and the enzymatic methods, the second strand of the cDNA was synthesized with a Klenow fragment using a primer corresponding to the 5'end of the ligated oligonucleotide described in Example 12.

Preferably, the primer is 20-25 bases in length. Methylated dCTP was also used for second strand synthesis in order to protect internal EcoRI sites in the cDNA from digestion during the cloning process.

Following cDNA synthesis, the cDNAs were cloned into pBlueScript as described in Example 15 below.

EXAMPLE 15

Insertion of cDNAs into BlueScript

Following second strand synthesis, the ends of the cDNA were blunted with T4 DNA polymerase (Biolabs) and the cDNA was digested with EcoRI. Since methylated dCTP was used during cDNA synthesis, the EcoRI site present in the tag was the only site which was hemi-methylated. Consequently, only the EcoRI site in the oligonucleotide tag was susceptible to EcoRI digestion. The cDNA was then size fractionated using exclusion chromatography (AcA, Biosepra). Fractions corresponding to cDNAs of more than 150 bp were pooled and ethanol precipitated. The cDNA was directionally cloned into the Smal and EcoRI ends of the phagemid pBlueScript vector (Stratagene). The ligation mixture was electroporated into bacteria and propagated under appropriate antibiotic selection.

Clones containing the oligonucleotide tag attached were selected as described in Example 16 below.

EXAMPLE 16

Selection of Clones Having the Oligonucleotide Tag Attached Thereto

The plasmid DNAs containing 5' EST libraries made as described above were purified (Qiagen). A positive selection of the tagged clones was performed as follows. Briefly, in this selection procedure, the plasmid DNA was converted to single stranded DNA using gene II endonuclease of the phage F1 in combination with an exonuclease (Chang et al., Gene 127:95-8, 1993) such as exonuclease III or T7 gene 6 exonuclease. The resulting single stranded DNA was then purified using paramagnetic beads as described by Fry et al., Biotechniques, 13: 124-131, 1992. In this procedure, the single stranded DNA was hybridized with a biotinylated oligonucleotide having a sequence corresponding to the 3' end of the oligonucleotide described in Example 13. Preferably, the primer has a length of 20-25 bases. Clones including a sequence complementary to the biotinylated oligonucleotide were captured by incubation with streptavidin coated magnetic beads followed by magnetic selection. After capture of the positive clones, the plasmid DNA was released from the magnetic beads and converted into double stranded DNA using a DNA polymerase such as the ThermoSequenase obtained from Amersham Pharmacia Biotech. Alternatively, protocols such as the Gene Trapper kit (Gibco BRL) may be used. The double stranded DNA was then electroporated into bacteria. The percentage of positive clones having the 5' tag oligonucleotide was estimated to typically rank between 90 and 98% using dot blot analysis.

Following electroporation, the libraries were ordered in 384-microtiter plates (MTP). A copy of the MTP was stored for future needs. Then the libraries were transferred into 96 MTP and sequenced as described below.

EXAMPLE 17

Sequencing of Inserts in Selected Clones

Plasmid inserts were first amplified by PCR on PE 9600 thermocyclers (Perkin-Elmer), using standard SETA-A and SETA-B primers (Genset SA), AmpliTaqGold (Perkin-Elmer), dNTPs (Boehringer), buffer and cycling conditions as recommended by the Perkin-Elmer Corporation.

PCR products were then sequenced using automatic ABI Prism 377 sequencers (Perkin Elmer, Applied Biosystems Division, Foster City, CA). Sequencing reactions were performed using PE 9600 thermocyclers (Perkin Elmer) with standard dye-primer chemistry and ThermoSequenase (Amersham Life Science). The primers used were either T7 or 21M13 (available from Genset SA) as appropriate. The primers were labeled with the JOE, FAM, ROX and TAMRA dyes. The dNTPs and ddNTPs used in the sequencing reactions were purchased from Boehringer. Sequencing buffer, reagent concentrations and cycling conditions were as recommended by Amersham.

Following the sequencing reaction, the samples were precipitated with EtOH, resuspended in formamide loading buffer, and loaded on a standard 4% acrylamide gel. Electrophoresis was performed for 2.5 hours at 3000V on an ABI 377 sequencer, and the sequence data were collected and analyzed using the ABI Prism DNA Sequencing 30 Analysis Software, version 2.1.2.

The sequence data from the 44 cDNA libraries made as described above were transferred to a proprietary database, where quality control and validation steps were performed. A proprietary base-caller ("Trace"), working using a Unix system automatically flagged suspect peaks, taking into account the shape of the peaks, the inter-peak resolution, and the noise level. The proprietary base-caller also performed an automatic trimming. Any stretch of 25 or

WO 99/31236 PCT/IB98/02122

fewer bases having more than 4 suspect peaks was considered unreliable and was discarded. Sequences corresponding to cloning vector or ligation oligonucleotides were automatically removed from the EST sequences. However, the resulting EST sequences may contain 1 to 5 bases belonging to the above mentioned sequences at their 5' end. If needed, these can easily be removed on a case by case basis.

Thereafter, the sequences were transferred to the proprietary NETGENE™ Database for further analysis as described below.

Following sequencing as described above, the sequences of the 5' ESTs were entered in a proprietary database called NETGENETM for storage and manipulation. It will be appreciated by those skilled in the art that the data could be stored and manipulated on any medium which can be read and accessed by a computer. Computer readable media include magnetically readable media, optically readable media, or electronically readable media. For example, the computer readable media may be a hard disc, a floppy disc, a magnetic tape, CD-ROM, RAM, or ROM as well as other types of other media known to those skilled in the art.

In addition, the sequence data may be stored and manipulated in a variety of data processor programs in a variety of formats. For example, the sequence data may be stored as text in a word processing file, such as

MicrosoftWORD or WORDPERFECT or as an ASCII file in a variety of database programs familiar to those of skill in the art, such as DB2, SYBASE, or ORACLE.

The computer readable media on which the sequence information is stored may be in a personal computer, a network, a server or other computer systems known to those skilled in the art. The computer or other system preferably includes the storage media described above, and a processor for accessing and manipulating the sequence data.

Once the sequence data has been stored it may be manipulated and searched to locate those stored sequences which contain a desired nucleic acid sequence or which encode a protein having a particular functional domain. For example, the stored sequence information may be compared to other known sequences to identify homologies, motifs implicated in biological function, or structural motifs.

20

Programs which may be used to search or compare the stored sequences include the MacPattern (EMBL),

25 BLAST, and BLAST2 program series (NCBI), basic local alignment search tool programs for nucleotide (BLASTN) and
peptide (BLASTX) comparisons (Altschul et al, J. Mol. Biol. 215: 403 (1990)) and FASTA (Pearson and Lipman, Proc.

Natl. Acad. Sci. USA, 85: 2444 (1988)). The BLAST programs then extend the alignments on the basis of defined
match and mismatch criteria.

Motifs which may be detected using the above programs include sequences encoding leucine zippers, helix-turn30 helix motifs, glycosylation sites, ubiquitination sites, alpha helices, and beta sheets, signal sequences encoding signal peptides which direct the secretion of the encoded proteins, sequences implicated in transcription regulation such as homeoboxes, acidic stretches, enzymatic active sites, substrate binding sites, and enzymatic cleavage sites.

25

Before searching the cDNAs in the NETGENE™ database for sequence motifs of interest, cDNAs derived from mRNAs which were not of interest were identified and eliminated from further consideration as described in Example 18 below.

EXAMPLE 18

Elimination of Undesired Sequences from Further Consideration

5' ESTs in the NETGENE™ database which were derived from undesired sequences such as transfer RNAs, ribosomal RNAs, mitochondrial RNAs, procaryotic RNAs, fungal RNAs, Alu sequences, L1 sequences, or repeat sequences were identified using the FASTA and BLASTN programs with the parameters listed in Table II.

To eliminate 5' ESTs encoding tRNAs from further consideration, the 5' EST sequences were compared to the 10 sequences of 1190 known tRNAs obtained from EMBL release 38, of which 100 were human. The comparison was performed using FASTA on both strands of the 5' ESTs. Sequences having more than 80% homology over more than 60 nucleotides were identified as tRNA. Of the 144,341 sequences screened, 26 were identified as tRNAs and eliminated from further consideration.

To eliminate 5' ESTs encoding rRNAs from further consideration, the 5' EST sequences were compared to the 15 sequences of 2497 known rRNAs obtained from EMBL release 38, of which 73 were human. The comparison was performed using BLASTN on both strands of the 5' ESTs with the parameter S = 108. Sequences having more than 80% homology over stretches longer than 40 nucleotides were identified as rRNAs. Of the 144,341 sequences screened, 3,312 were identified as rRNAs and eliminated from further consideration.

To eliminate 5' ESTs encoding mtRNAs from further consideration, the 5' EST sequences were compared to 20 the sequences of the two known mitochondrial genomes for which the entire genomic sequences are available and all sequences transcribed from these mitochondrial genomes including tRNAs, rRNAs, and mRNAs for a total of 38 sequences. The comparison was performed using BLASTN on both strands of the 5' ESTs with the parameter S=108. Sequences having more than 80% homology over stretches longer than 40 nucleotides were identified as mtRNAs. Of the 144,341 sequences screened, 6,110 were identified as mtRNAs and eliminated from further consideration.

Sequences which might have resulted from exogenous contaminants were eliminated from further consideration by comparing the 5' EST sequences to release 46 of the EMBL bacterial and fungal divisions using BLASTN with the parameter S = 144. All sequences having more than 90% homology over at least 40 nucleotides were identified as exogenous contaminants. Of the 42 cDNA libraries examined, the average percentages of procaryotic and fungal sequences contained therein were 0.2% and 0.5% respectively. Among these sequences, only one could be 30 identified as a sequence specific to fungi. The others were either fungal or procaryotic sequences having homologies with vertebrate sequences or including repeat sequences which had not been masked during the electronic comparison.

In addition, the 5' ESTs were compared to 6093 Alu sequences and 1115 L1 sequences to mask 5' ESTs containing such repeat sequences from further consideration. 5' ESTs including THE and MER repeats, SSTR sequences or satellite, micro-satellite, or telomeric repeats were also eliminated from further consideration. On average, 11.5% of

the sequences in the libraries contained repeat sequences. Of this 11.5%, 7% contained Alu repeats, 3.3% contained £1 repeats and the remaining 1.2% were derived from the other types of repetitive sequences which were screened. These percentages are consistent with those found in cDNA libraries prepared by other groups. For example, the cDNA libraries of Adams et al. contained between 0% and 7.4% Alu repeats depending on the source of the RNA which was used to prepare the cDNA library (Adams et al., *Nature* 377:174, 1996).

The sequences of those 5' ESTs remaining after the elimination of undesirable sequences were compared with the sequences of known human mRNAs to determine the accuracy of the sequencing procedures described above.

EXAMPLE 19

Measurement of Sequencing Accuracy by Comparison to Known Sequences

To further determine the accuracy of the sequencing procedure described above, the sequences of 5' ESTs derived from known sequences were identified and compared to the known sequences. First, a FASTA analysis with overhangs shorter than 5 bp on both ends was conducted on the 5' ESTs to identify those matching an entry in the public human mRNA database. The 6655 5' ESTs which matched a known human mRNA were then realigned with their cognate mRNA and dynamic programming was used to include substitutions, insertions, and deletions in the list of "errors" which would be recognized. Errors occurring in the last 10 bases of the 5' EST sequences were ignored to avoid the inclusion of spurious cloning sites in the analysis of sequencing accuracy.

This analysis revealed that the sequences incorporated in the NETGENE™ database had an accuracy of more than 99.5%.

To determine the efficiency with which the above selection procedures select cDNAs which include the 5' ends of their corresponding mRNAs, the following analysis was performed.

EXAMPLE 20

Determination of Efficiency of 5' EST Selection

To determine the efficiency at which the above selection procedures isolated 5' ESTs which included sequences close to the 5' end of the mRNAs from which they were derived, the sequences of the ends of the 5' ESTs which were derived from the elongation factor 1 subunit α and ferritin heavy chain genes were compared to the known cDNA sequences for these genes. Since the transcription start sites for the elongation factor 1 subunit α and ferritin heavy chain are well characterized, they may be used to determine the percentage of 5' ESTs derived from these genes which included the authentic transcription start sites.

For both genes, more than 95% of the cDNAs included sequences close to or upstream of the 5' end of the 30 corresponding mRNAs.

To extend the analysis of the reliability of the procedures for isolating 5' ESTs from ESTs in the NETGENETM database, a similar analysis was conducted using a database composed of human mRNA sequences extracted from GenBank database release 97 for comparison. For those 5' ESTs derived from mRNAs included in the GeneBank database, more than 85% had their 5' ends close to the 5' ends of the known sequence. As some of the mRNA

sequences available in the GenBank database are deduced from genomic sequences, a 5' end matching with these sequences will be counted as an internal match. Thus, the method used here underestimates the yield of ESTs including the authentic 5' ends of their corresponding mRNAs.

The EST libraries made above included multiple 5' ESTs derived from the same mRNA. The sequences of such 5' ESTs were compared to one another and the longest 5' ESTs for each mRNA were identified. Overlapping cDNAs were assembled into continuous sequences (contigs). The resulting continuous sequences were then compared to public databases to gauge their similarity to known sequences, as described in Example 21 below.

EXAMPLE 21

Clustering of the 5' ESTs and Calculation of Novelty Indices for cDNA Libraries

For each sequenced EST library, the sequences were clustered by the 5' end. Each sequence in the library was compared to the others with BLASTN2 (direct strand, parameters S = 107). ESTs with High Scoring Segment Pairs (HSPs) at least 25 bp long, having 95% identical bases and beginning closer than 10 bp from each EST 5' end were grouped. The longest sequence found in the cluster was used as representative of the cluster. A global clustering between libraries was then performed leading to the definition of super-contigs.

To assess the yield of new sequences within the EST libraries, a novelty rate (NR) was defined as: NR = 100 X (Number of new unique sequences found in the library/Total number of sequences from the library). Typically, novelty rating range between 10% and 41% depending on the tissue from which the EST library was obtained. For most of the libraries, the random sequencing of 5' EST libraries was pursued until the novelty rate reached 20%.

Following characterization as described above, the collection of 5' ESTs in NETGENETM was screened to 20 identify those 5' ESTs bearing potential signal sequences as described in Example 22 below.

EXAMPLE 22

Identification of Potential Signal Sequences in 5' ESTs

The 5' ESTs in the NETGENETM database were screened to identify those having an uninterrupted open reading frame (ORF) longer than 45 nucleotides beginning with an ATG codon and extending to the end of the EST.

25 Approximately half of the cDNA sequences in NETGENETM contained such an ORF. The ORFs of these 5' ESTs were searched to identify potential signal motifs using slight modifications of the procedures disclosed in Von Heijne, G. A New Method for Predicting Signal Sequence Cleavage Sites. Nucleic Acids Res. 14:4683-4690 (1986). Those 5' EST sequences encoding a 15 amino acid long stretch with a score of at least 3.5 in the Von Heijne signal peptide identification matrix were considered to possess a signal sequence. Those 5' ESTs which matched a known human mRNA or EST sequence and had a 5' end more than 20 nucleotides downstream of the known 5' end were excluded from further analysis. The remaining cDNAs having signal sequences therein were included in a database called SIGNALTAGTM.

To confirm the accuracy of the above method for identifying signal sequences, the analysis of Example 23 was performed.

-27-

EXAMPLE 23

Confirmation of Accuracy of Identification of Potential Signal Sequences in 5' ESTs

The accuracy of the above procedure for identifying signal sequences encoding signal peptides was evaluated by applying the method to the 43 amino terminal amino acids of all human SwissProt proteins. The computed Von Heijne score for each protein was compared with the known characterization of the protein as being a secreted protein or a non-secreted protein. In this manner, the number of non-secreted proteins having a score higher than 3.5 (false positives) and the number of secreted proteins having a score lower than 3.5 (false negatives) could be calculated.

Using the results of the above analysis, the probability that a peptide encoded by the 5' region of the mRNA is in fact a genuine signal peptide based on its Von Heijne's score was calculated based on either the assumption that 10% of human proteins are secreted or the assumption that 20% of human proteins are secreted. The results of this analysis are shown in Figures 2 and 3.

Using the above method of identifying secretory proteins, 5' ESTs for human glucagon, gamma interferon induced monokine precursor, secreted cyclophilin-like protein, human pleiotropin, and human biotinidase precursor all of which are polypeptides which are known to be secreted, were obtained. Thus, the above method successfully identified those 5' ESTs which encode a signal peptide.

To confirm that the signal peptide encoded by the 5' ESTs actually functions as a signal peptide, the signal sequences from the 5' ESTs may be cloned into a vector designed for the identification of signal peptides. Some signal peptide identification vectors are designed to confer the ability to grow in selective medium on host cells which have a signal sequence operably inserted into the vector. For example, to confirm that a 5' EST encodes a genuine signal peptide, the signal sequence of the 5' EST may be inserted upstream and in frame with a non-secreted form of the yeast invertase gene in signal peptide selection vectors such as those described in U.S. Patent No. 5,536,637. Growth of host cells containing signal sequence selection vectors having the signal sequence from the 5' EST inserted therein confirms that the 5' EST encodes a genuine signal peptide.

Alternatively, the presence of a signal peptide may be confirmed by cloning the extended cDNAs obtained using
the ESTs into expression vectors such as pXT1 (as described below), or by constructing promoter-signal sequencereporter gene vectors which encode fusion proteins between the signal peptide and an assayable reporter protein. After
introduction of these vectors into a suitable host cell, such as COS cells or NIH 3T3 cells, the growth medium may be
harvested and analyzed for the presence of the secreted protein. The medium from these cells is compared to the
medium from cells containing vectors lacking the signal sequence or extended cDNA insert to identify vectors which
encode a functional signal peptide or an authentic secreted protein.

Those 5' ESTs which encoded a signal peptide, as determined by the method of Example 22 above, were further grouped into four categories based on their homology to known sequences. The categorization of the 5' ESTs is described in Example 24 below.

25

30

Categorization of 5' ESTs Encoding a Signal Peptide

Those 5' ESTs having a sequence not matching any known vertebrate sequence nor any publicly available EST sequence were designated "new." Of the sequences in the SIGNALTAGTM database, 947 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those 5' ESTs having a sequence not matching any vertebrate sequence but matching a publicly known EST were designated "EST-ext", provided that the known EST sequence was extended by at least 40 nucleotides in the 5' direction. Of the sequences in the SIGNALTAGTM database, 150 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those ESTs not matching any vertebrate sequence but matching a publicly known EST without extending the known EST by at least 40 nucleotides in the 5' direction were designated "EST." Of the sequences in the SIGNALTAGTM database, 599 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those 5' ESTs matching a human mRNA sequence but extending the known sequence by at least 40 nucleotides in the 5' direction were designated "VERT-ext." Of the sequences in the SIGNALTAGTM database, 23 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category. Included in this category was a 5' EST which extended the known sequence of the human translocase mRNA by more than 200 bases in the 5' direction. A 5' EST which extended the sequence of a human tumor suppressor gene in the 5' direction was also identified.

Figure 4 shows the distribution of 5' ESTs in each category and the number of 5' ESTs in each category having a given minimum von Heijne's score.

Each of the 5' ESTs was categorized based on the tissue from which its corresponding mRNA was obtained, 20 as described below in Example 25.

EXAMPLE 25

Categorization of Expression Patterns

Figure 5 shows the tissues from which the mRNAs corresponding to the 5' ESTs in each of the above described categories were obtained.

In addition to categorizing the 5' ESTs by the tissue from which the cDNA library in which they were first identified was obtained, the spatial and temporal expression patterns of the mRNAs corresponding to the 5' ESTs, as well as their expression levels, may be determined as described in Example 26 below. Characterization of the spatial and temporal expression patterns and expression levels of these mRNAs is useful for constructing expression vectors capable of producing a desired level of gene product in a desired spatial or temporal manner, as will be discussed in more detail below.

In addition, 5' ESTs whose corresponding mRNAs are associated with disease states may also be identified. For example, a particular disease may result from lack of expression, over expression, or under expression of an mRNA corresponding to a 5' EST. By comparing mRNA expression patterns and quantities in samples taken from healthy

individuals with those from individuals suffering from a particular disease, 5' ESTs responsible for the disease may be identified.

It will be appreciated that the results of the above characterization procedures for 5' ESTs also apply to extended cDNAs (obtainable as described below) which contain sequences adjacent to the 5' ESTs. It will also be appreciated that if it is desired to defer characterization until extended cDNAs have been obtained rather than characterizing the ESTs themselves, the above characterization procedures can be applied to characterize the extended cDNAs after their isolation.

EXAMPLE 26

Evaluation of Expression Levels and Patterns of mRNAs

Corresponding to 5' ESTs or Extended cDNAs

10

Expression levels and patterns of mRNAs corresponding to 5' ESTs or extended cDNAs (obtainable as described below) may be analyzed by solution hybridization with long probes as described in International Patent Application No. WO 97/05277. Briefly, a 5' EST, extended cDNA, or fragment thereof corresponding to the gene encoding the mRNA to be characterized is inserted at a cloning site immediately downstream of a bacteriophage (T3, T7 or SP6) RNA polymerase promoter to produce antisense RNA. Preferably, the 5' EST or extended cDNA has 100 or more nucleotides. The plasmid is linearized and transcribed in the presence of ribonucleotides comprising modified ribonucleotides (i.e. biotin-UTP and DIG-UTP). An excess of this doubly labeled RNA is hybridized in solution with mRNA isolated from cells or tissues of interest. The hybridizations are performed under standard stringent conditions (40-50°C for 16 hours in an 80% formamide, 0.4 M NaCl buffer, pH 7-8). The unhybridized probe is removed by digestion with ribonucleases specific for single-stranded RNA (i.e. RNases CL3, T1, Phy M, U2 or A). The presence of the biotin-UTP modification enables capture of the hybrid on a microtitration plate coated with streptavidin. The presence of the DIG modification enables the hybrid to be detected and quantified by ELISA using an anti-DIG antibody coupled to alkaline phosphatase.

The 5' ESTs, extended cDNAs, or fragments thereof may also be tagged with nucleotide sequences for the serial analysis of gene expression (SAGE) as disclosed in UK Patent Application No. 2 305 241 A. In this method, cDNAs are prepared from a cell, tissue, organism or other source of nucleic acid for which it is desired to determine gene expression patterns. The resulting cDNAs are separated into two pools. The cDNAs in each pool are cleaved with a first restriction endonuclease, called an "anchoring enzyme," having a recognition site which is likely to be present at least once in most cDNAs. The fragments which contain the 5' or 3' most region of the cleaved cDNA are isolated by binding to a capture medium such as streptavidin coated beads. A first oligonucleotide linker having a first sequence for hybridization of an amplification primer and an internal restriction site for a "tagging endonuclease" is ligated to the digested cDNAs in the first pool. Digestion with the second endonuclease produces short "tag" fragments from the cDNAs.

25

A second oligonucleotide having a second sequence for hybridization of an amplification primer and an internal restriction site is ligated to the digested cDNAs in the second pool. The cDNA fragments in the second pool are also digested with the "tagging endonuclease" to generate short "tag" fragments derived from the cDNAs in the second pool. The "tags" resulting from digestion of the first and second pools with the anchoring enzyme and the tagging 5 endonuclease are ligated to one another to produce "ditags." In some embodiments, the ditags are concatamerized to produce ligation products containing from 2 to 200 ditags. The tag sequences are then determined and compared to the sequences of the 5' ESTs or extended cDNAs to determine which 5' ESTs or extended cDNAs are expressed in the cell, tissue, organism, or other source of nucleic acids from which the tags were derived. In this way, the expression pattern of the 5' ESTs or extended cDNAs in the cell, tissue, organism, or other source of nucleic acids is obtained.

Quantitative analysis of gene expression may also be performed using arrays. As used herein, the term array means a one dimensional, two dimensional, or multidimensional arrangement of full length cDNAs (i.e. extended cDNAs which include the coding sequence for the signal peptide, the coding sequence for the mature protein, and a stop codon), extended cDNAs, 5' ESTs or fragments of the full length cDNAs, extended cDNAs, or 5' ESTs of sufficient length to permit specific detection of gene expression. Preferably, the fragments are at least 15 nucleotides in length. More 15 preferably, the fragments are at least 100 nucleotides in length. More preferably, the fragments are more than 100 nucleotides in length. In some embodiments the fragments may be more than 500 nucleotides in length.

For example, quantitative analysis of gene expression may be performed with full length cDNAs, extended cDNAs, 5' ESTs, or fragments thereof in a complementary DNA microarray as described by Schena et al. (Science 270:467-470, 1995; Proc. Natl. Acad. Sci. U.S.A. 93:10614-10619, 1996). Full length cDNAs, extended cDNAs, 5' 20 ESTs or fragments thereof are amplified by PCR and arrayed from 96-well microtiter plates onto silylated microscope slides using high-speed robotics. Printed arrays are incubated in a humid chamber to allow rehydration of the array elements and rinsed, once in 0.2% SDS for 1 min, twice in water for 1 min and once for 5 min in sodium borohydride solution. The arrays are submerged in water for 2 min at 95°C, transferred into 0.2% SDS for 1 min, rinsed twice with water, air dried and stored in the dark at 25°C.

Cell or tissue mRNA is isolated or commercially obtained and probes are prepared by a single round of reverse transcription. Probes are hybridized to 1 cm² microarrays under a 14 x 14 mm glass coverslip for 6-12 hours at 60°C. Arrays are washed for 5 min at 25°C in low stringency wash buffer (1 x SSC/0.2% SDS), then for 10 min at room temperature in high stringency wash buffer (0.1 x SSC/0.2% SDS). Arrays are scanned in 0.1 x SSC using a fluorescence laser scanning device fitted with a custom filter set. Accurate differential expression measurements are 30 obtained by taking the average of the ratios of two independent hybridizations.

Quantitative analysis of the expression of genes may also be performed with full length cDNAs, extended cDNAs, 5' ESTs, or fragments thereof in complementary DNA arrays as described by Pietu et al. (Genome Research 6:492-503, 1996). The full length cDNAs, extended cDNAs, 5' ESTs or fragments thereof are PCR amplified and spotted on membranes. Then, mRNAs originating from various tissues or cells are labeled with radioactive nucleotides.

After hybridization and washing in controlled conditions, the hybridized mRNAs are detected by phospho-imaging or autoradiography. Duplicate experiments are performed and a quantitative analysis of differentially expressed mRNAs is then performed.

Alternatively, expression analysis of the 5' ESTs or extended cDNAs can be done through high density nucleotide arrays as described by Lockhart et al. (Nature Biotechnology 14: 1675-1680, 1996) and Sosnowsky et al. (Proc. Natl. Acad. Sci. 94:1119-1123, 1997). Oligonucleotides of 15-50 nucleotides corresponding to sequences of the 5' ESTs or extended cDNAs are synthesized directly on the chip (Lockhart et al., *supra*) or synthesized and then addressed to the chip (Sosnowski et al., *supra*). Preferably, the oligonucleotides are about 20 nucleotides in length.

cDNA probes labeled with an appropriate compound, such as biotin, digoxigenin or fluorescent dye, are
synthesized from the appropriate mRNA population and then randomly fragmented to an average size of 50 to 100 nucleotides. The said probes are then hybridized to the chip. After washing as described in Lockhart et al., *supra* and application of different electric fields (Sosnowsky et al., Proc. Natl. Acad. Sci. 94:1119-1123)., the dyes or labeling compounds are detected and quantified. Duplicate hybridizations are performed. Comparative analysis of the intensity of the signal originating from cDNA probes on the same target oligonucleotide in different cDNA samples indicates a differential expression of the mRNA corresponding to the 5' EST or extended cDNA from which the oligonucleotide sequence has been designed.

III. Use of 5' ESTs to Clone Extended cDNAs and to Clone the Corresponding Genomic DNAs

Once 5' ESTs which include the 5' end of the corresponding mRNAs have been selected using the procedures described above, they can be utilized to isolate extended cDNAs which contain sequences adjacent to the 5' ESTs. The extended cDNAs may include the entire coding sequence of the protein encoded by the corresponding mRNA, including the authentic translation start site, the signal sequence, and the sequence encoding the mature protein remaining after cleavage of the signal peptide. Such extended cDNAs are referred to herein as "full length cDNAs." Alternatively, the extended cDNAs may include only the sequence encoding the mature protein remaining after cleavage of the signal peptide, or only the sequence encoding the signal peptide.

Example 27 below describes a general method for obtaining extended cDNAs. Example 28 below describes the cloning and sequencing of several extended cDNAs, including extended cDNAs which include the entire coding sequence and authentic 5' end of the corresponding mRNA for several secreted proteins.

The methods of Examples 27, 28, and 29 can also be used to obtain extended cDNAs which encode less than the entire coding sequence of the secreted proteins encoded by the genes corresponding to the 5' ESTs. In some embodiments, the extended cDNAs isolated using these methods encode at least 10 amino acids of one of the proteins encoded by the sequences of SEQ ID NOs: 40-140 and 242-377. In further embodiments, the extended cDNAs encode at least 20 amino acids of the proteins encoded by the sequences of SEQ ID NOs: 40-140 and 242-377. In further embodiments, the extended cDNAs encode at least 30 amino amino acids of the sequences of SEQ ID NOs: 40-140 and

242-377. In a preferred embodiment, the extended cDNAs encode a full length protein sequence, which includes the protein coding sequences of SEQ ID NOs: 40-140 and 242-377.

EXAMPLE 27

General Method for Using 5' ESTs to Clone and Sequence Extended cDNAs

The following general method has been used to quickly and efficiently isolate extended cDNAs including sequence adjacent to the sequences of the 5' ESTs used to obtain them. This method may be applied to obtain extended cDNAs for any 5' EST in the NETGENETM database, including those 5' ESTs encoding secreted proteins. The method is summarized in Figure 6.

1. Obtaining Extended cDNAs

10 a) First strand synthesis

5

The method takes advantage of the known 5' sequence of the mRNA. A reverse transcription reaction is conducted on purified mRNA with a poly 14dT primer containing a 49 nucleotide sequence at its 5' end allowing the addition of a known sequence at the end of the cDNA which corresponds to the 3' end of the mRNA. For example, the primer may have the following sequence: 5'-ATC GTT GAG ACT CGT ACC AGC AGA GTC ACG AGA GAG ACT ACA CGG TAC TGG TTT TTT TTT TTT TTVN -3' (SEQ ID NO:14). Those skilled in the art will appreciate that other sequences may also be added to the poly dT sequence and used to prime the first strand synthesis. Using this primer and a reverse transcriptase such as the Superscript II (Gibco BRL) or Rnase H Minus M-MLV (Promega) enzyme, a reverse transcript anchored at the 3' polyA site of the RNAs is generated.

After removal of the mRNA hybridized to the first cDNA strand by alkaline hydrolysis, the products of the alkaline hydrolysis and the residual poly dT primer are eliminated with an exclusion column such as an AcA34 (Biosepra) matrix as explained in Example 11.

b) Second strand synthesis

30

A pair of nested primers on each end is designed based on the known 5' sequence from the 5' EST and the known 3' end added by the poly dT primer used in the first strand synthesis. Software used to design primers are either based on GC content and melting temperatures of oligonucleotides, such as OSP (Illier and Green, *PCR Meth. Appl.* 1:124-128, 1991), or based on the octamer frequency disparity method (Griffais et al., *Nucleic Acids Res.* 19: 3887-3891, 1991 such as PC-Rare (http://bioinformatics.weizmann.ac.il/software/PC-Rare/doc/manuel.html).

Preferably, the nested primers at the 5' end are separated from one another by four to nine bases. The 5' primer sequences may be selected to have melting temperatures and specificities suitable for use in PCR.

Preferably, the nested primers at the 3' end are separated from one another by four to nine bases. For example, the nested 3' primers may have the following sequences: (5'- CCA GCA GAG TCA CGA GAG AGA CTA CAC GG -3' (SEQ ID NO:15), and 5'- CAC GAG AGA GAC TAC ACG GTA CTG G -3' (SEQ ID NO:16). These primers were selected because they have melting temperatures and specificities compatible with their use in PCR. However, those skilled in the art will appreciate that other sequences may also be used as primers.

The first PCR run of 25 cycles is performed using the Advantage Tth Polymerase Mix (Clontech) and the outerprimer from each of the nested pairs. A second 20 cycle PCR using the same enzyme and the inner primer from each of the nested pairs is then performed on 1/2500 of the first PCR product. Thereafter, the primers and nucleotides are removed.

5 2. Seguencing of Full Length Extended cDNAs or Fragments Thereof

Due to the lack of position constraints on the design of 5' nested primers compatible for PCR use using the OSP software, amplicons of two types are obtained. Preferably, the second 5' primer is located upstream of the translation initiation codon thus yielding a nested PCR product containing the whole coding sequence. Such a full length extended cDNA undergoes a direct cloning procedure as described in section a below. However, in some cases, the second 5' primer is located downstream of the translation initiation codon, thereby yielding a PCR product containing only part of the ORF. Such incomplete PCR products are submitted to a modified procedure described in section b below.

a) Nested PCR products containing complete ORFs

When the resulting nested PCR product contains the complete coding sequence, as predicted from the 5'EST sequence, it is closed in an appropriate vector such as pED6dpc2, as described in section 3.

b) Nested PCR products containing incomplete ORFs

When the amplicon does not contain the complete coding sequence, intermediate steps are necessary to obtain both the complete coding sequence and a PCR product containing the full coding sequence. The complete coding sequence can be assembled from several partial sequences determined directly from different PCR products as described in the following section.

Once the full coding sequence has been completely determined, new primers compatible for PCR use are designed to obtain amplicons containing the whole coding region. However, in such cases, 3' primers compatible for PCR use are located inside the 3' UTR of the corresponding mRNA, thus yielding amplicons which lack part of this region, i.e. the polyA tract and sometimes the polyadenylation signal, as illustrated in figure 6. Such full length extended cDNAs are then cloned into an appropriate vector as described in section 3.

c) Sequencing extended cDNAs

Sequencing of extended cDNAs is performed using a Die Terminator approach with the AmpliTaq DNA polymerase FS kit available from Perkin Elmer.

In order to sequence PCR fragments, primer walking is performed using software such as OSP to choose

30 primers and automated computer software such as ASMG (Sutton et al., *Genome Science Technol.* 1: 9-19, 1995) to construct contigs of walking sequences including the initial 5' tag using minimum overlaps of 32 nucleotides. Preferably, primer walking is performed until the sequences of full length cDNAs are obtained.

Completion of the sequencing of a given extended cDNA fragment is assessed as follows. Since sequences located after a polyA tract are difficult to determine precisely in the case of uncloned products, sequencing and primer

walking processes for PCR products are interrupted when a polyA tract is identified in extended cDNAs obtained as described in case b. The sequence length is compared to the size of the nested PCR product obtained as described above. Due to the limited accuracy of the determination of the PCR product size by gel electrophoresis, a sequence is considered complete if the size of the obtained sequence is at least 70 % the size of the first nested PCR product. If the 5 length of the sequence determined from the computer analysis is not at least 70% of the length of the nested PCR product, these PCR products are cloned and the sequence of the insertion is determined. When Northern blot data are available, the size of the mRNA detected for a given PCR product is used to finally assess that the sequence is complete. Sequences which do not fulfill the above criteria are discarded and will undergo a new isolation procedure.

Sequence data of all extended cDNAs are then transferred to a proprietary database, where quality controls 10 and validation steps are carried out as described in example 15.

3. Cloning of Full Length Extended cDNAs

20

The PCR product containing the full coding sequence is then cloned in an appropriate vector. For example, the extended cDNAs can be cloned into the expression vector pED6dpc2 (DiscoverEase, Genetics Institute, Cambridge, MA) as follows. The structure of pED6dpc2 is shown in Figure 7. pED6dpc2 vector DNA is prepared with blunt ends by 15 performing an EcoRI digestion followed by a fill in reaction. The blunt ended vector is dephosphorylated. After removal of PCR primers and ethanol precipitation, the PCR product containing the full coding sequence or the extended cDNA obtained as described above is phosphorylated with a kinase subsequently removed by phenol-Sevag extraction and precipitation. The double stranded extended cDNA is then ligated to the vector and the resulting expression plasmid introduced into appropriate host cells.

Since the PCR products obtained as described above are blunt ended molecules that can be cloned in either direction, the orientation of several clones for each PCR product is determined. Then, 4 to 10 clones are ordered in microtiter plates and subjected to a PCR reaction using a first primer located in the vector close to the cloning site and a second primer located in the portion of the extended cDNA corresponding to the 3' end of the mRNA. This second primer may be the antisense primer used in anchored PCR in the case of direct cloning (case a) or the antisense primer located 25 inside the 3'UTR in the case of indirect cloning (case b). Clones in which the start codon of the extended cDNA is operably linked to the promoter in the vector so as to permit expression of the protein encoded by the extended cDNA are conserved and sequenced. In addition to the ends of cDNA inserts, approximately 50 bp of vector DNA on each side of the cDNA insert are also sequenced.

The cloned PCR products are then entirely sequenced according to the aforementioned procedure. In this case, 30 contig assembly of long fragments is then performed on walking sequences that have already contigated for uncloned PCR products during primer walking. Sequencing of cloned amplicons is complete when the resulting contigs include the whole coding region as well as overlapping sequences with vector DNA on both ends.

4. Computer Analysis of Full Length Extended cDNA

Sequences of all full length extended cDNAs are then submitted to further analysis as described below and using the parameters found in Table II with the following modifications. For screening of miscellaneous subdivisions of Genbank, FASTA was used instead of BLASTN and 15 nucleotide of homology was the limit instead of 17. For Alu detection, BLASTN was used with the following parameters: S = 72; identity = 70%; and length = 40 nucleotides.

- Polyadenylation signal and polyA tail which were not search for the 5' ESTs were searched. For polyadenylation signal detection the signal (AATAAA) was searched with one permissible mismatch in the last ten nucleotides preceding the 5' end of the polyA. For the polyA, a stretch of 8 amino acids in the last 20 nucleotides of the sequence was searched with BLAST2N in the sense strand with the following parameters (W = 6, S = 10, E = 1000, and identity = 90%). Finally, patented sequences and ORF homologies were searched using, respectively, BLASTN and BLASTP on GenSEQ
- 10 (Derwent's database of patented nucleotide sequences) and SWISSPROT for ORFs with the following parameters (W = 8 and B = 10). Before examining the extended full length cDNAs for sequences of interest, extended cDNAs which are not of interest are searched as follows.

a) Elimination of undesired sequences

Although 5'ESTs were checked to remove contaminant sequences as described in Example 18, a last verification was carried out to identify extended cDNAs sequences derived from undesired sequences such as vector RNAs, transfer RNAs, ribosomal rRNAs, mitochondrial RNAs, prokaryotic RNAs and fungal RNAs using the FASTA and BLASTN programs on both strands of extended cDNAs as described below.

To identify the extended cDNAs encoding vector RNAs, extended cDNAs are compared to the known sequences of vector RNA using the FASTA program. Sequences of extended cDNAs with more than 90% homology over stretches of 15 nucleotides are identified as vector RNA.

To identify the extended cDNAs encoding tRNAs, extended cDNA sequences were compared to the sequences of 1190 known tRNAs obtained from EMBL release 38, of which 100 were human. Sequences of extended cDNAs having more than 80% homology over 60 nucleotides using FASTA were identified as tRNA.

To identify the extended cDNAs encoding rRNAs, extended cDNA sequences were compared to the sequences
of 2497 known rRNAs obtained from EMBL release 38, of which 73 were human. Sequences of extended cDNAs having
more than 80% homology over stretches longer than 40 nucleotides using BLASTN were identified as rRNAs.

To identify the extended cDNAs encoding mtRNAs, extended cDNA sequences were compared to the sequences of the two known mitochondrial genomes for which the entire genomic sequences are available and all sequences transcribed from these mitochondrial genomes including tRNAs, rRNAs, and mRNAs for a total of 38 sequences. Sequences of extended cDNAs having more than 80% homology over stretches longer than 40 nucleotides using BLASTN were identified as mtRNAs.

Sequences which might have resulted from other exogenous contaminants were identified by comparing extended cDNA sequences to release 105 of Genbank bacterial and fungal divisions. Sequences of extended cDNAs

having more than 90% homology over 40 nucleotides using BLASTN were identified as exogenous prokaryotic or fungal contaminants.

In addition, extended cDNAs were searched for different repeat sequences, including Alu sequences, L1 sequences, THE and MER repeats, SSTR sequences or satellite, micro-satellite, or telomeric repeats. Sequences of extended cDNAs with more than 70% homology over 40 nucleotide stretches using BLASTN were identified as repeat sequences and masked in further identification procedures. In addition, clones showing extensive homology to repeats, i.e., matches of either more than 50 nucleotides if the homology was at least 75% or more than 40 nucleotides if the homology was at least 90%, were flagged.

b) Identification of structural features

10

20

25

Structural features, e.g. polyA tail and polyadenylation signal, of the sequences of full length extended cDNAs are subsequently determined as follows.

A polyA tail is defined as a homopolymeric stretch of at least 11 A with at most one alternative base within it.

The polyA tail search is restricted to the last 20 nt of the sequence and limited to stretches of 11 consecutive A's because sequencing reactions are often not readable after such a polyA stretch. Stretches with 100% homology over 6 nucleotides are identified as polyA tails.

To search for a polyadenylation signal, the polyA tail is clipped from the full-length sequence. The 50 bp preceding the polyA tail are searched for the canonic polyadenylation AAUAAA signal allowing one mismatch to account for possible sequencing errors and known variation in the canonical sequence of the polyadenylation signal.

c) Identification of functional features

Functional features, e.g. ORFs and signal sequences, of the sequences of full length extended cDNAs were subsequently determined as follows.

The 3 upper strand frames of extended cDNAs are searched for ORFs defined as the maximum length fragments beginning with a translation initiation codon and ending with a stop codon. ORFs encoding at least 20 amino acids are preferred.

Each found ORF is then scanned for the presence of a signal peptide in the first 50 amino-acids or, where appropriate, within shorter regions down to 20 amino acids or less in the ORF, using the matrix method of von Heijne (Nuc. Acids Res. 14: 4683-4690 (1986)) and the modification described in Example 22.

d) Homology to either nucleotidic or proteic sequences

Sequences of full length extended cDNAs are then compared to known sequences on a nucleotidic or proteic 30 basis.

Sequences of full length extended cDNAs are compared to the following known nucleic acid sequences: vertebrate sequences (Genbank), EST sequences (Genbank), patented sequences (Geneseqn) and recently identified sequences (Genbank daily releases) available at the time of filing for the priority documents. Full length cDNA sequences are also compared to the sequences of a private database (Genset internal sequences) in order to find sequences that

have already been identified by applicants. Sequences of full length extended cDNAs with more than 90% homology over 30 nucleotides using either BLASTN or BLAST2N as indicated in Table III are identified as sequences that have already been described. Matching vertebrate sequences are subsequently examined using FASTA; full length extended cDNAs with more than 70% homology over 30 nucleotides are identified as sequences that have already been described.

ORFs encoded by full length extended cDNAs as defined in section c) are subsequently compared to known amino acid sequences found in Swissprot release CHP, PIR release PIR# and Genpept release GPEPT public databases using BLASTP with the parameter W = 8 and allowing a maximum of 10 matches. Sequences of full length extended cDNAs showing extensive homology to known protein sequences are recognized as already identified proteins.

In addition, the three-frame conceptual translation products of the top strand of full length extended cDNAs

are compared to publicly known amino acid sequences of Swissprot using BLASTX with the parameter E=0.001.

Sequences of full length extended cDNAs with more than 70% homology over 30 amino acid stretches are detected as already identified proteins.

5. Selection of Cloned Full Length Sequences of the Present Invention

Cloned full length extended cDNA sequences that have already been characterized by the aforementioned computer analysis are then submitted to an automatic procedure in order to preselect full length extended cDNAs containing sequences of interest.

a) Automatic sequence preselection

All complete cloned full length extended cDNAs clipped for vector on both ends are considered. First, a negative selection is operated in order to eliminate unwanted cloned sequences resulting from either contaminants or PCR artifacts as follows. Sequences matching contaminant sequences such as vector RNA, tRNA, mtRNA, rRNA sequences are discarded as well as those encoding ORF sequences exhibiting extensive homology to repeats as defined in section 4 a). Sequences obtained by direct cloning using nested primers on 5' and 3' tags (section 1. case a) but lacking polyA tail are discarded. Only ORFs containing a signal peptide and ending either before the polyA tail (case a) or before the end of the cloned 3'UTR (case b) are kept. Then, ORFs containing unlikely mature proteins such as mature proteins which size is less than 20 amino acids or less than 25% of the immature protein size are eliminated.

In the selection of the OFR, priority was given to the ORF and the frame corresponding to the polypeptides described in SignalTag Patents (United States Patent Application Serial Nos: 08/905,223; 08/905,135; 08/905,051; 08/905,144; 08/905,279; 08/904,468; 08/905,134; and 08/905,133). If the ORF was not found among the OFRs described in the SignalTag Patents, the ORF encoding the signal peptide with the highest score according to Von Heijne method as defined in Example 22 was chosen. If the scores were identical, then the longest ORF was chosen.

Sequences of full length extended cDNA clones are then compared pairwise with BLAST after masking of the repeat sequences. Sequences containing at least 90% homology over 30 nucleotides are clustered in the same class. Each cluster is then subjected to a cluster analysis that detects sequences resulting from internal priming or from

alternative splicing, identical sequences or sequences with several frameshifts. This automatic analysis serves as a basis for manual selection of the sequences.

b) Manual sequence selection

30

Manual selection is carried out using automatically generated reports for each sequenced full length extended cDNA clone. During this manual procedures, a selection is operated between clones belonging to the same class as follows. ORF sequences encoded by clones belonging to the same class are aligned and compared. If the homology between nucleotidic sequences of clones belonging to the same class is more than 90% over 30 nucleotide stretches or if the homology between amino acid sequences of clones belonging to the same class is more than 80% over 20 amino acid stretches, than the clones are considered as being identical. The chosen ORF is the best one according to the criteria mentioned below. If the nucleotide and amino acid homologies are less than 90% and 80% respectively, the clones are said to encode distinct proteins which can be both selected if they contain sequences of interest.

Selection of full length extended cDNA clones encoding sequences of interest is performed using the following criteria. Structural parameters (initial tag, polyadenylation site and signal) are first checked. Then, homologies with known nucleic acids and proteins are examined in order to determine whether the clone sequence match a known nucleic/proteic sequence and, in the latter case, its covering rate and the date at which the sequence became public. If there is no extensive match with sequences other than ESTs or genomic DNA, or if the clone sequence brings substantial new information, such as encoding a protein resulting from alternative slicing of an mRNA coding for an already known protein, the sequence is kept. Examples of such cloned full length extended cDNAs containing sequences of interest are described in Example 28. Sequences resulting from chimera or double inserts as assessed by homology to other

EXAMPLE 28

Cloning and Sequencing of Extended cDNAs

The procedure described in Example 27 above was used to obtain the extended cDNAs of the present invention. Using this approach, the full length cDNA of SEQ ID NO:17 was obtained. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide MKKVLLLITAILAVAVG (SEQ ID NO: 18) having a von Heijne score of 8.2.

The full length cDNA of SEQ ID NO:19 was also obtained using this procedure. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide MWWFQQGLSFLPSALVIWTSA (SEQ ID NO:20) having a von Heijne score of 5.5.

Another full length cDNA obtained using the procedure described above has the sequence of SEQ ID NO:21.

This cDNA, falls into the "EST-ext" category described above and encodes the signal peptide

MVLTTLPSANSANSPVNMPTTGPNSLSYASSALSPCLT (SEQ ID NO:22) having a von Heijne score of 5.9.

PCT/IB98/02122

10

30

The above procedure was also used to obtain a full length cDNA having the sequence of SEQ ID NO:23. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide ILSTVTALTFAXA (SEQ ID NO:24) having a von Heijne score of 5.5.

The full length cDNA of SEQ ID NO:25 was also obtained using this procedure. This cDNA falls into the "new" category described above and encodes a signal peptide LVLTLCTLPLAVA (SEQ ID NO:26) having a von Heijne score of 10.1.

The full length cDNA of SEQ ID NO:27 was also obtained using this procedure. This cDNA falls into the "new" category described above and encodes a signal peptide LWLLFFLVTAIHA (SEQ ID NO:28) having a von Heijne score of 10.7.

The above procedures were also used to obtain the extended cDNAs of the present invention. 5' ESTs expressed in a variety of tissues were obtained as described above. The appended sequence listing provides the tissues from which the extended cDNAs were obtained. It will be appreciated that the extended cDNAs may also be expressed in tissues other than the tissue listed in the sequence listing.

5' ESTs obtained as described above were used to obtain extended cDNAs having the sequences of SEQ ID

NOs: 40-140 and 242-377. Table IV provides the sequence identification numbers of the extended cDNAs of the present invention, the locations of the full coding sequences in SEQ ID NOs: 40-140 and 242-377 (i.e. the nucleotides encoding both the signal peptide and the mature protein, listed under the heading FCS location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the signal peptides (listed under the heading SigPep Location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the mature proteins generated by cleavage of the signal peptides (listed under the heading Mature Polypeptide Location in Table IV), the locations in SEQ ID NOs: 40-140 and 242-377 of stop codons (listed under the heading Stop Codon Location in Table IV), the locations in SEQ ID NOs: 40-140 and 242-377 of polyA signals (listed under the heading Poly A Signal Location in Table IV) and the locations of polyA sites (listed under the heading Poly A Site Location in Table IV).

The polypeptides encoded by the extended cDNAs were screened for the presence of known structural or functional motifs or for the presence of signatures, small amino acid sequences which are well conserved amongst the members of a protein family. The conserved regions have been used to derive consensus patterns or matrices included in the PROSITE data bank, in particular in the file prosite.dat (Release 13.0 of November 1995, located at http://expasy.hcuge.ch/sprot/prosite.html. Prosite_convert and prosite_scan programs (http://ulrec3.unil.ch/ftpserveur/prosite_scan) were used to find signatures on the extended cDNAs.

For each pattern obtained with the prosite_convert program from the prosite.dat file, the accuracy of the detection on a new protein sequence has been tested by evaluating the frequency of irrelevant hits on the population of human secreted proteins included in the data bank SWISSPROT. The ratio between the number of hits on shuffled proteins (with a window size of 20 amino acids) and the number of hits on native (unshuffled) proteins was used as an index. Every pattern for which the ration was greater than 20% (one hit on shuffled proteins for 5 hits on native

proteins) was skipped during the search with prosite_scan. The program used to shuffle protein sequences (db_shuffled) and the program used to determine the statistics for each pattern in the protein data banks (prosite_statistics) are available on the ftp site http://ulrec3.unil.ch/ftpserveur/prosite_scan.

Table V lists the sequence identification numbers of the polypeptides of SEQ ID NOs: 141-241 and 378-513, the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the full length polypeptide (second column), the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the signal peptides (third column), and the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the mature polypeptide created by cleaving the signal peptide from the full length polypeptide (fourth column).

The nucleotide sequences of the sequences of SEQ ID Nos: 40-140 and 242-377 and the amino acid sequences of SEQ ID NOs: 40-140 and 242-377 (i.e. amino acid sequences of SEQ ID NOs: 141-241 and 378-513) are provided in the appended sequence listing. In some instances, the sequences are preliminary and may include some incorrect or ambiguous sequences or amino acids. The sequences of SEQ ID NOs: 40-140 and 242-377 can readily be screened for any errors therein and any sequence ambiguities can be resolved by resequencing a fragment containing such errors or ambiguities on both strands. Nucleic acid fragments for resolving sequencing errors or ambiguities may be obtained from the deposited clones or can be isolated using the techniques described herein. Resolution of any such ambiguities or errors may be facilitated by using primers which hybridize to sequences located close to the ambiguous or erroneous sequences. For example, the primers may hybridize to sequences within 50-75 bases of the ambiguity or error. Upon resolution of an error or ambiguity, the corresponding corrections can be made in the protein sequences encoded by the DNA containing the error or ambiguity. For example, in the sequences of the present invention, ambiguities in the sequence of SEQ ID NO: 131 were resolved. The amino acid sequence of the protein encoded by a particular clone can also be determined by expression of the clone in a suitable host cell, collecting the protein, and determining its sequence.

For each amino acid sequence, Applicants have identified what they have determined to be the reading frame best identifiable with sequence information available at the time of filing. Some of the amino acid sequences may contain "Xaa" designators. These "Xaa" designators indicate either (1) a residue which cannot be identified because of nucleotide sequence ambiguity or (2) a stop codon in the determined sequence where Applicants believe one should not exist (if the sequence were determined more accurately).

Cells containing the extended cDNAs (SEQ ID NOs: 40-140 and 242-377) of the present invention in the vector pED6dpc2, are maintained in permanent deposit by the inventors at Genset, S.A., 24 Rue Royale, 75008 Paris, France.

Pools of cells containing the extended cDNAs (SEQ ID NOs: 40-140 and 242-377), from which cells containing a particular polynucleotide are obtainable, were deposited with the American Type Culture Collection, 10801 University Blvd., Manassas, VA 20110-2209 or the European Collection of Cell Cultures, Vaccine Research and Production Laboratory, Public Health Laboratory Service, Centre for Applied Microbiology and Research, Porton Down, Salisbury, Wiltshire SP4 OJG, United Kingdom. Each extended cDNA clone has been transfected into separate bacterial cells (E-

20

coli) for this composite deposit. Table VI lists the deposit numbers of the clones containing the extended cDNAs of the present invention. Table VII provides the internal designation number assigned to each SEQ ID NO and indicates whether the sequence is a nucleic acid sequence or a protein sequence.

Each extended cDNA can be removed from the pED6dpc2 vector in which it was deposited by performing a 5 Notl, Pstl double digestion to produce the appropriate fragment for each clone. The proteins encoded by the extended cDNAs may also be expressed from the promoter in pED6dpc2.

Bacterial cells containing a particular clone can be obtained from the composite deposit as follows:

An oligonucleotide probe or probes should be designed to the sequence that is known for that particular clone. This sequence can be derived from the sequences provided herein, or from a combination of those sequences. The design 10 of the oligonucleotide probe should preferably follow these parameters:

- (a) It should be designed to an area of the sequence which has the fewest ambiguous bases ("N's"), if any;
- (b) Preferably, the probe is designed tc have a T_m of approx. 80°C (assuming 2 degrees for each A or T and 4 degrees for each G or C). However, probes having melting temperatures between 40 °C and 80 °C may also be used provided that specificity is not lost.

The oligonucleotide should preferably be labeled with (-[32P]ATP (specific activity 6000 Ci/mmole) and T4 polynucleotide kinase using commonly employed techniques for labeling oligonucleotides. Other labeling techniques can also be used. Unincorporated label should preferably be removed by gel filtration chromatography or other established methods. The amount of radioactivity incorporated into the probe should be quantified by measurement in a scintillation counter. Preferably, specific activity of the resulting probe should be approximately 4X106 dpm/pmole.

The bacterial culture containing the pool of full-length clones should preferably be thawed and 100 µl of the stock used to inoculate a sterile culture flask containing 25 ml of sterile L-broth containing ampicillin at 100 ug/ml. The culture should preferably be grown to saturation at 37°C, and the saturated culture should preferably be diluted in fresh L-broth. Aliquots of these dilutions should preferably be plated to determine the dilution and volume which will yield approximately 5000 distinct and well-separated colonies on solid bacteriological media containing L-broth containing 25 ampicillin at 100 μg/ml and agar at 1.5% in a 150 mm petri dish when grown overnight at 37°C. Other known methods of obtaining distinct, well-separated colonies can also be employed.

Standard colony hybridization procedures should then be used to transfer the colonies to nitrocellulose filters and lyse, denature and bake them.

The filter is then preferably incubated at 65°C for 1 hour with gentle agitation in 6X SSC (20X stock is 30 175.3 g NaC1/liter, 88.2 g Na citrate/liter, adjusted to pH 7.0 with NaOH) containing 0.5% SDS, 100 pg/ml of yeast RNA, and 10 mM EDTA (approximately 10 mL per 150 mm filter). Preferably, the probe is then added to the hybridization mix at a concentration greater than or equal to 1X10⁶ dpm/mL. The filter is then preferably incubated at 65°C with gentle agitation overnight. The filter is then preferably washed in 500 mL of 2X SSC/0.1% SDS at room temperature with gentle shaking for 15 minutes. A third wash with 0.1X SSC/0.5% SDS at 65°C for 30 minutes to

1 hour is optional. The filter is then preferably dried and subjected to autoradiography for sufficient time to visualize the positives on the X-ray film. Other known hybridization methods can also be employed.

The positive colonies are picked, grown in culture, and plasmid DNA isolated using standard procedures. The clones can then be verified by restriction analysis, hybridization analysis, or DNA sequencing.

The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR can be done with primers designed at both ends of the extended cDNA insertion. For example, a PCR reaction may be conducted using a primer having the sequence GGCCATACACTTGAGTGAC (SEQ ID NO:38) and a primer having the sequence ATATAGACAAACGCACACC (SEQ. ID. NO:39). The PCR product which corresponds to the extended cDNA can then be manipulated using standard cloning 10 techniques familiar to those skilled in the art.

In addition to PCR based methods for obtaining extended cDNAs, traditional hybridization based methods may also be employed. These methods may also be used to obtain the genomic DNAs which encode the mRNAs from which the 5' ESTs were derived, mRNAs corresponding to the extended cDNAs, or nucleic acids which are homologous to extended cDNAs or 5' ESTs. Example 29 below provides an example of such methods.

15

30

5

EXAMPLE 29

Methods for Obtaining Extended cDNAs or Nucleic Acids Homologous to Extended cDNAs or 5' ESTs

A full length cDNA library can be made using the strategies described in Examples 13, 14, 15, and 16 above by replacing the random nonamer used in Example 14 with an oligo-dT primer. For instance, the oligonucleotide of SEQ ID 20 N0:14 may be used.

Alternatively, a cDNA library or genomic DNA library may be obtained from a commercial source or made using techniques familiar to those skilled in the art. The library includes cDNAs which are derived from the mRNA corresponding to a 5' EST or which have homology to an extended cDNA or 5' EST. The cDNA library or genomic DNA library is hybridized to a detectable probe comprising at least 10 consecutive nucleotides from the 5' EST or extended 25 cDNA using conventional techniques. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST or extended cDNA. More preferably, the probe comprises at least 20-30 consecutive nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises at least 30 nucleotides from the 5' EST or extended cDNA. In other embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the 5' EST or extended cDNA.

Techniques for identifying cDNA clones in a cDNA library which hybridize to a given probe sequence are disclosed in Sambrook et al., Molecular Cloning: A Laboratory Manual 2d Ed., Cold Spring Harbor Laboratory Press, 1989. The same techniques may be used to isolate genomic DNAs.

Briefly, cDNA or genomic DNA clones which hybridize to the detectable probe are identified and isolated for further manipulation as follows. A probe comprising at least 10 consecutive nucleotides from the 5' EST or extended cDNA is labeled with a detectable label such as a radioisotope or a fluorescent molecule. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST or extended cDNA. More preferably, the probe comprises 20-30 consecutive nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises more than 30 nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the 5' EST or extended cDNA.

Techniques for labeling the probe are well known and include phosphorylation with polynucleotide kinase, nick translation, in vitro transcription, and non-radioactive techniques. The cDNAs or genomic DNAs in the library are transferred to a nitrocellulose or nylon filter and denatured. After incubation of the filter with a blocking solution, the filter is contacted with the labeled probe and incubated for a sufficient amount of time for the probe to hybridize to cDNAs or genomic DNAs containing a sequence capable of hybridizing to the probe.

By varying the stringency of the hybridization conditions used to identify extended cDNAs or genomic DNAs which hybridize to the detectable probe, extended cDNAS having different levels of homology to the probe can be identified and isolated. To identify extended cDNAs or genomic DNAs having a high degree of homology to the probe sequence, the melting temperature of the probe may be calculated using the following formulas:

For probes between 14 and 70 nucleotides in length the melting temperature (Tm) is calculated using the formula: Tm = 81.5 + 16.6(log [Na +]) + 0.41(fraction G + C)-(600/N) where N is the length of the probe.

If the hybridization is carried out in a solution containing formamide, the melting temperature may be calculated using the equation Tm = 81.5 + 16.6(log [Na +]) + 0.41(fraction G + C)-(0.63% formamide)-(600/N) where N is the length of the probe.

Prehybridization may be carried out in 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100µg denatured fragmented salmon sperm DNA or 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100µg denatured fragmented salmon sperm DNA, 50% formamide. The formulas for SSC and Denhardt's solutions are listed in Sambrook et al., supra.

Hybridization is conducted by adding the detectable probe to the prehybridization solutions listed above. Where
the probe comprises double stranded DNA, it is denatured before addition to the hybridization solution. The filter is
contacted with the hybridization solution for a sufficient period of time to allow the probe to hybridize to extended
cDNAs or genomic DNAs containing sequences complementary thereto or homologous thereto. For probes over 200
nucleotides in length, the hybridization may be carried out at 15-25°C below the Tm. For shorter probes, such as
oligonucleotide probes, the hybridization may be conducted at 15-25°C below the Tm. Preferably, for hybridizations in
6X SSC, the hybridization is conducted at approximately 68°C. Preferably, for hybridizations in 50% formamide
containing solutions, the hybridization is conducted at approximately 42°C.

All of the foregoing hybridizations would be considered to be under "stringent" conditions. Following hybridization, the filter is washed in 2X SSC, 0.1% SDS at room temperature for 15 minutes. The filter is then washed

with 0.1X SSC, 0.5% SDS at room temperature for 30 minutes to 1 hour. Thereafter, the solution is washed at the hybridization temperature in 0.1X SSC, 0.5% SDS. A final wash is conducted in 0.1X SSC at room temperature.

Extended cDNAs, nucleic acids homologous to extended cDNAs or 5' ESTs, or genomic DNAs which have hybridized to the probe are identified by autoradiography or other conventional techniques.

The above procedure may be modified to identify extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs having decreasing levels of homology to the probe sequence. For example, to obtain extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs of decreasing homology to the detectable probe, less stringent conditions may be used. For example, the hybridization temperature may be decreased in increments of 5°C from 68°C to 42°C in a hybridization buffer having a Na+ concentration of approximately 1M. Following 10 hybridization, the filter may be washed with 2X SSC, 0.5% SDS at the temperature of hybridization. These conditions are considered to be "moderate" conditions above 50°C and "low" conditions below 50°C.

Alternatively, the hybridization may be carried out in buffers, such as 6X SSC, containing formamide at a temperature of 42°C. In this case, the concentration of formamide in the hybridization buffer may be reduced in 5% increments from 50% to 0% to identify clones having decreasing levels of homology to the probe. Following 15 hybridization, the filter may be washed with 6X SSC, 0.5% SDS at 50°C. These conditions are considered to be "moderate" conditions above 25% formamide and "low" conditions below 25% formamide.

Extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs which have hybridized to the probe are identified by autoradiography.

If it is desired to obtain nucleic acids homologous to extended cDNAs, such as allelic variants thereof or nucleic 20 acids encoding proteins related to the proteins encoded by the extended cDNAs, the level of homology between the hybridized nucleic acid and the extended cDNA or 5' EST used as the probe may readily be determined. To determine the level of homology between the hybridized nucleic acid and the extended cDNA or 5'EST from which the probe was derived, the nucleotide sequences of the hybridized nucleic acid and the extended cDNA or 5'EST from which the probe was derived are compared. For example, using the above methods, nucleic acids having at least 95% nucleic acid 25 homology to the extended cDNA or 5'EST from which the probe was derived may be obtained and identified. Similarly, by using progressively less stringent hybridization conditions one can obtain and identify nucleic acids having at least 90%, at least 85%, at least 80% or at least 75% homology to the extended cDNA or 5'EST from which the probe was derived. The level of homology between the hybridized nucleic acid and the extended cDNA or 5' EST used as the probe may be further determined using BLAST2N; parameters may be adapted depending on the sequence length and degree of 30 homology studied. In such comparisons, the default parameters or the parameters listed in Tables II and III may be used.

To determine whether a clone encodes a protein having a given amount of homology to the protein encoded by the extended cDNA or 5' EST, the amino acid sequence encoded by the extended cDNA or 5' EST is compared to the amino acid sequence encoded by the hybridizing nucleic acid. Homology is determined to exist when an amino acid sequence in the extended cDNA or 5' EST is closely related to an amino acid sequence in the hybridizing nucleic acid. A

sequence is closely related when it is identical to that of the extended cDNA or 5' EST or when it contains one or more amino acid substitutions therein in which amino acids having similar characteristics have been substituted for one another. Using the above methods, one can obtain nucleic acids encoding proteins having at least 95%, at least 90%, at least 85%, at least 80% or at least 75% homology to the proteins encoded by the extended cDNA or 5'EST from which the probe was derived. Using the above methods and algorithms such as FASTA with parameters depending on the sequence length and degree of homology studied the level of homology may be determined. In determining the level of homology using FASTA, the default parameters or the parameters listed in Tables II or III may be used.

Alternatively, extended cDNAs may be prepared by obtaining mRNA from the tissue, cell, or organism of interest using mRNA preparation procedures utilizing poly A selection procedures or other techniques known to those skilled in the art. A first primer capable of hybridizing to the poly A tail of the mRNA is hybridized to the mRNA and a reverse transcription reaction is performed to generate a first cDNA strand.

The first cDNA strand is hybridized to a second primer containing at least 10 consecutive nucleotides of the sequences of the 5' EST for which an extended cDNA is desired. Preferably, the primer comprises at least 12, 15, or 17 consecutive nucleotides from the sequences of the 5' EST. More preferably, the primer comprises 20-30 consecutive nucleotides from the sequences of the 5' EST. In some embodiments, the primer comprises more than 30 nucleotides from the sequences of the 5' EST. If it is desired to obtain extended cDNAs containing the full protein coding sequence, including the authentic translation initiation site, the second primer used contains sequences located upstream of the translation initiation site. The second primer is extended to generate a second cDNA strand complementary to the first cDNA strand. Alternatively, RTPCR may be performed as described above using primers from both ends of the cDNA to be obtained.

Extended cDNAs containing 5' fragments of the mRNA may be prepared by contacting an mRNA comprising the sequence of the 5' EST for which an extended cDNA is desired with a primer comprising at least 10 consecutive nucleotides of the sequences complementary to the 5' EST, hybridizing the primer to the mRNAs, and reverse transcribing the hybridized primer to make a first cDNA strand from the mRNAs. Preferably, the primer comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST. More preferably, the primer comprises 20-30 consecutive nucleotides from the 5' EST.

Thereafter, a second cDNA strand complementary to the first cDNA strand is synthesized. The second cDNA strand may be made by hybridizing a primer complementary to sequences in the first cDNA strand to the first cDNA strand and extending the primer to generate the second cDNA strand.

The double stranded extended cDNAs made using the methods described above are isolated and cloned. The extended cDNAs may be cloned into vectors such as plasmids or viral vectors capable of replicating in an appropriate host cell. For example, the host cell may be a bacterial, mammalian, avian, or insect cell.

30

Techniques for isolating mRNA, reverse transcribing a primer hybridized to mRNA to generate a first cDNA strand, extending a primer to make a second cDNA strand complementary to the first cDNA strand, isolating the double

30

stranded cDNA and cloning the double stranded cDNA are well known to those skilled in the art and are described in Current Protocols in Molecular Biology, John Wiley 503 Sons, Inc. 1997 and Sambrook et al. Molecular Cloning: A Laboratory Manual, Second Edition, Cold Spring Harbor Laboratory Press, 1989.

Alternatively, kits for obtaining full length cDNAs, such as the GeneTrapper (Cat. No. 10356-020, Gibco, BRL),
may be used for obtaining full length cDNAs or extended cDNAs. In this approach, full length or extended cDNAs are
prepared from mRNA and cloned into double stranded phagemids. The cDNA library in the double stranded phagemids is
then rendered single stranded by treatment with an endonuclease, such as the Gene II product of the phage F1, and
Exonuclease III as described in the manual accompanying the GeneTrapper kit. A biotinylated oligonucleotide comprising
the sequence of a 5' EST, or a fragment containing at least 10 nucleotides thereof, is hybridized to the single stranded
phagemids. Preferably, the fragment comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST. More
preferably, the fragment comprises 20-30 consecutive nucleotides from the 5' EST. In some procedures, the fragment
may comprise more than 30 consecutive nucleotides from the 5' EST. For example, the fragment may comprises at least
40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the 5' EST.

Hybrids between the biotinylated oligonucleotide and phagemids having inserts containing the 5' EST sequence are isolated by incubating the hybrids with streptavidin coated paramagnetic beads and retrieving the beads with a magnet. Thereafter, the resulting phagemids containing the 5' EST sequence are released from the beads and converted into double stranded DNA using a primer specific for the 5' EST sequence. The resulting double stranded DNA is transformed into bacteria. Extended cDNAs containing the 5' EST sequence are identified by colony PCR or colony hybridization.

A plurality of extended cDNAs containing full length protein coding sequences or sequences encoding only the mature protein remaining after the signal peptide is cleaved may be provided as cDNA libraries for subsequent evaluation of the encoded proteins or use in diagnostic assays as described below.

IV. Expression of Proteins Encoded by Extended cDNAs Isolated Using 5' ESTs

Extended cDNAs containing the full protein coding sequences of their corresponding mRNAs or portions

thereof, such as cDNAs encoding the mature protein, may be used to express the secreted proteins or portions thereof which they encode as described in Example 30 below. If desired, the extended cDNAs may contain the sequences encoding the signal peptide to facilitate secretion of the expressed protein. It will be appreciated that a plurality of extended cDNAs containing the full protein coding sequences or portions thereof may be simultaneously cloned into expression vectors to create an expression library for analysis of the encoded proteins as described below.

EXAMPLE 30

Expression of the Proteins Encoded by Extended cDNAs or Portions Thereof

To express the proteins encoded by the extended cDNAs or portions thereof, nucleic acids containing the coding sequence for the proteins or portions thereof to be expressed are obtained as described in Examples 27-29 and cloned into a suitable expression vector. If desired, the nucleic acids may contain the sequences encoding the signal

PCT/IB98/02122 WO 99/31236

-47-

peptide to facilitate secretion of the expressed protein. For example, the nucleic acid may comprise the sequence of one of SEQ ID NOs: 40-140 and 242-377 listed in Table IV and in the accompanying sequence listing. Alternatively, the nucleic acid may comprise those nucleotides which make up the full coding sequence of one of the sequences of SEQ ID NOs: 40-140 and 242-377 as defined in Table IV above.

5

20

It will be appreciated that should the extent of the full coding sequence (i.e. the sequence encoding the signal peptide and the mature protein resulting from cleavage of the signal peptide) differ from that listed in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, post-translational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the extent of the full coding sequences in the sequences of SEQ ID NOs. 40-140 and 242-377. 10 For example, the sequence of SEQ ID NO: 115 represents an alternatively spliced transcript of a previously identified mRNA.. Accordingly, the scope of any claims herein relating to nucleic acids containing the full coding sequence of one of SEQ ID NOs. 40-140 and 242-377 is not to be construed as excluding any readily identifiable variations from or equivalents to the full coding sequences listed in Table IV Similarly, should the extent of the full length polypeptides differ from those indicated in Table V as a result of any of the preceding factors, the scope of claims relating to polypeptides 15 comprising the amino acid sequence of the full length polypeptides is not to be construed as excluding any readily identifiable variations from or equivalents to the sequences listed in Table V.

Alternatively, the nucleic acid used to express the protein or portion thereof may comprise those nucleotides which encode the mature protein (i.e. the protein created by cleaving the signal peptide off) encoded by one of the sequences of SEQ ID NOs: 40-140 and 242-377 as defined in Table IV above.

It will be appreciated that should the extent of the sequence encoding the mature protein differ from that listed in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, posttranslational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the extent of the sequence encoding the mature protein in the sequences of SEQ ID NOs. 40-140 and 242-377. Accordingly, the scope of any claims herein relating to nucleic acids 25 containing the sequence encoding the mature protein encoded by one of SEQ ID Nos. 40-140 and 242-377 is not to be construed as excluding any readily identifiable variations from or equivalents to the sequences listed in Table IV. Thus, claims relating to nucleic acids containing the sequence encoding the mature protein encompass equivalents to the sequences listed in Table IV, such as sequences encoding biologically active proteins resulting from post-translational modification, enzymatic cleavage, or other readily identifiable variations from or equivalents to the secreted proteins in 30 addition to cleavage of the signal peptide. Similarly, should the extent of the mature polypeptides differ from those indicated in Table V as a result of any of the preceding factors, the scope of claims relating to polypeptides comprising the sequence of a mature protein included in the sequence of one of SEQ ID NOs. 141-241 and 378-513 is not to be construed as excluding any readily identifiable variations from or equivalents to the sequences listed in Table V. Thus, claims relating to polypeptides comprising the sequence of the mature protein encompass equivalents to the sequences

listed in Table IV, such as biologically active proteins resulting from post-translational modification, enzymatic cleavage, or other readily identifiable variations from or equivalents to the secreted proteins in addition to cleavage of the signal peptide. It will also be appreciated that should the biologically active form of the polypeptides included in the sequence of one of SEQ ID NOs. 141-241 and 378-513 or the nucleic acids encoding the biologically active form of the polypeptides differ from those identified as the mature polypeptide in Table V or the nucleotides encoding the mature polypeptide in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, post-translational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the amino acids in the biologically active form of the polypeptides and the nucleic acids encoding the biologically active form of the polypeptides. In such instances, the claims relating to polypetides comprising the mature protein included in one of SEQ ID NOs. 141-241 and 378-513 or nucleic acids comprising the nucleotides of one of SEQ ID NOs. 40-140 and 242-377 encoding the mature protein shall not be construed to exclude any readily identifiable variations from the sequences listed in Table IV and Table V.

In some embodiments, the nucleic acid used to express the protein or portion thereof may comprise those nucleotides which encode the signal peptide encoded by one of the sequences of SEO ID NOs: 40-140 and 242-377 as defined in Table IV above.

It will be appreciated that should the extent of the sequence encoding the signal peptide differ from that listed in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, post-translational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the extent of the sequence encoding the signal peptide in the sequences of SEQ ID NOs. 40-140 and 242-377. Accordingly, the scope of any claims herein relating to nucleic acids containing the sequence encoding the signal peptide encoded by one of SEQ ID Nos. 40-140 and 242-377 is not to be construed as excluding any readily identifiable variations from the sequences listed in Table IV. Similarly, should the extent of the signal peptides differ from those indicated in Table V as a result of any of the preceding factors, the scope of claims relating to polypeptides comprising the sequence of a signal peptide included in the sequence of one of SEQ ID NOs. 141-241 and 378-513 is not to be construed as excluding any readily identifiable variations from the sequences listed in Table V.

Alternatively, the nucleic acid may encode a polypeptide comprising at least 10 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In some embodiments, the nucleic acid may encode a polypeptide comprising at least 15 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-30. In other embodiments, the nucleic acid may encode a polypeptide comprising at least 25 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In other embodiments, the nucleic acid may encode a polypeptide comprising at least 60, at least 75, at least 100 or more than 100 consecutive amino acids of one of the sequences of SEQ ID Nos: 141-241 and 378-513.

The nucleic acids inserted into the expression vectors may also contain sequences upstream of the sequences encoding the signal peptide, such as sequences which regulate expression levels or sequences which confer tissue specific expression.

The nucleic acid encoding the protein or polypeptide to be expressed is operably linked to a promoter in an expression vector using conventional cloning technology. The expression vector may be any of the mammalian, yeast, insect or bacterial expression systems known in the art. Commercially available vectors and expression systems are available from a variety of suppliers including Genetics Institute (Cambridge, MA), Stratagene (La Jolla, California), Promega (Madison, Wisconsin), and Invitrogen (San Diego, California). If desired, to enhance expression and facilitate proper protein folding, the codon context and codon pairing of the sequence may be optimized for the particular expression organism in which the expression vector is introduced, as explained by Hatfield, et al., U.S. Patent No. 5,082,767.

The following is provided as one exemplary method to express the proteins encoded by the extended cDNAs corresponding to the 5' ESTs or the nucleic acids described above. First, the methionine initiation codon for the gene and the poly A signal of the gene are identified. If the nucleic acid encoding the polypeptide to be expressed lacks a methionine to serve as the initiation site, an initiating methionine can be introduced next to the first codon of the nucleic acid using conventional techniques. Similarly, if the extended cDNA lacks a poly A signal, this sequence can be added to the construct by, for example, splicing out the Poly A signal from pSG5 (Stratagene) using Bgll and Sall restriction endonuclease enzymes and incorporating it into the mammalian expression vector pXT1 (Stratagene). pXT1 contains the LTRs and a portion of the gag gene from Moloney Murine Leukemia Virus. The position of the LTRs in the construct allow efficient stable transfection. The vector includes the Herpes Simplex Thymidine Kinase promoter and the selectable neomycin gene. The extended cDNA or portion thereof encoding the polypeptide to be expressed is obtained by PCR from the bacterial vector using oligonucleotide primers complementary to the extended cDNA or portion thereof and containing restriction endonuclease sequences for Pst I incorporated into the 5' primer and Bglll at the 5' end of the corresponding cDNA 3' primer, taking care to ensure that the extended cDNA is positioned in frame with the poly A signal. The purified fragment obtained from the resulting PCR reaction is digested with PstI, blunt ended with an exonuclease, digested with Bgll II, purified and ligated to pXT1, now containing a poly A signal and digested with BgllI.

The ligated product is transfected into mouse NIH 3T3 cells using Lipofectin (Life Technologies, Inc., Grand Island, New York) under conditions outlined in the product specification. Positive transfectants are selected after growing the transfected cells in 600ug/ml G418 (Sigma, St. Louis, Missouri). Preferably the expressed protein is released into the culture medium, thereby facilitating purification.

Alternatively, the extended cDNAs may be cloned into pED6dpc2 as described above. The resulting pED6dpc2 constructs may be transfected into a suitable host cell, such as COS 1 cells. Methotrexate resistant cells are selected and expanded. Preferably, the protein expressed from the extended cDNA is released into the culture medium thereby facilitating purification.

WO 99/31236 PCT/IB98/02122

-50-

Proteins in the culture medium are separated by gel electrophoresis. If desired, the proteins may be ammonium sulfate precipitated or separated based on size or charge prior to electrophoresis.

As a control, the expression vector lacking a cDNA insert is introduced into host cells or organisms and the proteins in the medium are harvested. The secreted proteins present in the medium are detected using techniques such as Coomassie or silver staining or using antibodies against the protein encoded by the extended cDNA. Coomassie and silver staining techniques are familiar to those skilled in the art.

Antibodies capable of specifically recognizing the protein of interest may be generated using synthetic 15-mer peptides having a sequence encoded by the appropriate 5' EST, extended cDNA, or portion thereof. The synthetic peptides are injected into mice to generate antibody to the polypeptide encoded by the 5' EST, extended cDNA, or portion thereof.

Secreted proteins from the host cells or organisms containing an expression vector which contains the extended cDNA derived from a 5' EST or a portion thereof are compared to those from the control cells or organism. The presence of a band in the medium from the cells containing the expression vector which is absent in the medium from the control cells indicates that the extended cDNA encodes a secreted protein. Generally, the band corresponding to the protein encoded by the extended cDNA will have a mobility near that expected based on the number of amino acids in the open reading frame of the extended cDNA. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

Alternatively, if the protein expressed from the above expression vectors does not contain sequences directing its secretion, the proteins expressed from host cells containing an expression vector containing an insert encoding a secreted protein or portion thereof can be compared to the proteins expressed in host cells containing the expression vector without an insert. The presence of a band in samples from cells containing the expression vector with an insert which is absent in samples from cells containing the expression vector without an insert indicates that the desired protein or portion thereof is being expressed. Generally, the band will have the mobility expected for the secreted protein or portion thereof. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

The protein encoded by the extended cDNA may be purified using standard immunochromatography techniques. In such procedures, a solution containing the secreted protein, such as the culture medium or a cell extract, is applied to a column having antibodies against the secreted protein attached to the chromatography matrix. The secreted protein is allowed to bind the immunochromatography column. Thereafter, the column is washed to remove non-specifically bound proteins. The specifically bound secreted protein is then released from the column and recovered using standard techniques.

If antibody production is not possible, the extended cDNA sequence or portion thereof may be incorporated into expression vectors designed for use in purification schemes employing chimeric polypeptides. In such strategies the coding sequence of the extended cDNA or portion thereof is inserted in frame with the gene encoding the other half of

the chimera. The other half of the chimera may be β-globin or a nickel binding polypeptide encoding sequence. A chromatography matrix having antibody to β-globin or nickel attached thereto is then used to purify the chimeric protein. Protease cleavage sites may be engineered between the β-globin gene or the nickel binding polypeptide and the extended cDNA or portion thereof. Thus, the two polypeptides of the chimera may be separated from one another by protease digestion.

One useful expression vector for generating β-globin chimerics is pSG5 (Stratagene), which encodes rabbit β-globin. Intron II of the rabbit β-globin gene facilitates splicing of the expressed transcript, and the polyadenylation signal incorporated into the construct increases the level of expression. These techniques as described are well known to those skilled in the art of molecular biology. Standard methods are published in methods texts such as Davis et al.,

10 (Basic Methods in Molecular Biology, L.G. Davis, M.D. Dibner, and J.F. Battey, ed., Elsevier Press, NY, 1986) and many of the methods are available from Stratagene, Life Technologies, Inc., or Promega. Polypeptide may additionally be produced from the construct using in vitro translation systems such as the In vitro ExpressTM Translation Kit (Stratagene).

Following expression and purification of the secreted proteins encoded by the 5' ESTs, extended cDNAs, or fragments thereof, the purified proteins may be tested for the ability to bind to the surface of various cell types as described in Example 31 below. It will be appreciated that a plurality of proteins expressed from these cDNAs may be included in a panel of proteins to be simultaneously evaluated for the activities specifically described below, as well as other biological roles for which assays for determining activity are available.

EXAMPLE 31

20

Analysis of Secreted Proteins to Determine Whether they Bind to the Cell Surface

The proteins encoded by the 5' ESTs, extended cDNAs, or fragments thereof are cloned into expression vectors such as those described in Example 30. The proteins are purified by size, charge, immunochromatography or other techniques familiar to those skilled in the art. Following purification, the proteins are labeled using techniques known to those skilled in the art. The labeled proteins are incubated with cells or cell lines derived from a variety of organs or tissues to allow the proteins to bind to any receptor present on the cell surface. Following the incubation, the cells are washed to remove non-specifically bound protein. The labeled proteins are detected by autoradiography. Alternatively, unlabeled proteins may be incubated with the cells and detected with antibodies having a detectable label, such as a fluorescent molecule, attached thereto.

Specificity of cell surface binding may be analyzed by conducting a competition analysis in which various

amounts of unlabeled protein are incubated along with the labeled protein. The amount of labeled protein bound to the

cell surface decreases as the amount of competitive unlabeled protein increases. As a control, various amounts of an

unlabeled protein unrelated to the labeled protein is included in some binding reactions. The amount of labeled protein

bound to the cell surface does not decrease in binding reactions containing increasing amounts of unrelated unlabeled

protein, indicating that the protein encoded by the cDNA binds specifically to the cell surface.

As discussed above, secreted proteins have been shown to have a number of important physiological effects and, consequently, represent a valuable therapeutic resource. The secreted proteins encoded by the extended cDNAs or portions thereof made according to Examples 27-29 may be evaluated to determine their physiological activities as described below.

5 EXAMPLE 32

Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Cytokine, Cell Proliferation or Cell Differentiation Activity

As discussed above, secreted proteins may act as cytokines or may affect cellular proliferation or differentiation. Many protein factors discovered to date, including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein of the present invention is evidenced by any one of a number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10, B5, B9/11, BaF3, MC9/G, M+ (preB M+), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7c and CMK. The proteins encoded by the above extended cDNAs or portions thereof may be evaluated for their ability to regulate T cell or thymocyte proliferation in assays such as those described above or in the following references: Current Protocols in Immunology, Ed. by J.E. Coligan et al., Greene Publishing Associates and Wiley-Interscience; Takai et al. J. Immunol. 137:3494-3500, 1986. Bertagnolli et al. J. Immunol. 145:1706-1712, 1990. Bertagnolli et al., Cellular Immunology 133:327-341, 1991. Bertagnolli, et al. J. Immunol. 149:3778-3783, 1992; Bowman et al., J. Immunol. 152:1756-1761, 1994.

In addition, numerous assays for cytokine production and/or the proliferation of spleen cells, lymph node cells
and thymocytes are known. These include the techniques disclosed in Current Protocols in Immunology. J.E. Coligan
et al. Eds., Vol 1 pp. 3.12.1-3.12.14 John Wiley and Sons, Toronto. 1994; and Schreiber, R.D. Current Protocols in
Immunology., supra Vol 1 pp. 6.8.1-6.8.8, John Wiley and Sons, Toronto. 1994.

The proteins encoded by the cDNAs may also be assayed for the ability to regulate the proliferation and differentiation of hematopoietic or lymphopoietic cells. Many assays for such activity are familiar to those skilled in the art, including the assays in the following references: Bottomly, K., Davis, L.S. and Lipsky, P.E., Measurement of Human and Murine Interleukin 2 and Interleukin 4, Current Protocols in Immunology., J.E. Coligan et al. Eds. Vol 1 pp. 6.3.1-6.3.12, John Wiley and Sons, Toronto. 1991; deVries et al., J. Exp. Med. 173:1205-1211, 1991; Moreau et al., Nature 36:690-692, 1988; Greenberger et al., Proc. Natl. Acad. Sci. U.S.A. 80:2931-2938, 1983; Nordan, R., Measurement of Mouse and Human Interleukin 6 Current Protocols in Immunology. J.E. Coligan et al. Eds. Vol 1 pp. 6.6.1-6.6.5, John Wiley and Sons, Toronto. 1991; Smith et al., Proc. Natl. Acad. Sci. U.S.A. 83:1857-1861, 1986; Bennett, F., Giannotti, J., Clark, S.C. and Turner, K.J., Measurement of Human Interleukin 11 Current Protocols in Immunology. J.E. Coligan et al. Eds. Vol 1 pp. 6.15.1 John Wiley and Sons, Toronto. 1991; Ciarletta, A., Giannotti, J., Clark, S.C. and Turner, K.J., Measurement of Mouse and Human Interleukin 9 Current Protocols in Immunology. J.E. Coligan et al., Eds. Vol 1 pp. 6.13.1, John Wiley and Sons, Toronto. 1991.

The proteins encoded by the cDNAs may also be assayed for their ability to regulate T-cell responses to antigens. Many assays for such activity are familiar to those skilled in the art, including the assays described in the following references: Chapter 3 (In Vitro Assays for Mouse Lymphocyte Function), Chapter 6 (Cytokines and Their Cellular Receptors) and Chapter 7, (Immunologic Studies in Humans) in Current Protocols in Immunology, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Interscience; Weinberger et al., Proc. Natl. Acad. Sci. USA 77:6091-6095, 1980; Weinberger et al., Eur. J. Immun. 11:405-411, 1981; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988.

Those proteins which exhibit cytokine, cell proliferation, or cell differentiation activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which induction of cell proliferation or differentiation is

10 beneficial. Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

EXAMPLE 33

Assaying the Proteins Expressed from Extended cDNAs or Portions

Thereof for Activity as Immune System Regulators

The proteins encoded by the cDNAs may also be evaluated for their effects as immune regulators. For example, the proteins may be evaluated for their activity to influence thymocyte or splenocyte cytotoxicity. Numerous assays for such activity are familiar to those skilled in the art including the assays described in the following references: Chapter 3 (In Vitro Assays for Mouse Lymphocyte Function 3.1-3.19) and Chapter 7 (Immunologic studies in Humans) in 20 Current Protocols in Immunology, J.E. Coligan et al. Eds, Greene Publishing Associates and Wiley-Interscience; Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Bowman et al., J. Virology 61:1992-1998; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Brown et al., J. Immunol. 153:3079-3092, 1994.

The proteins encoded by the cDNAs may also be evaluated for their effects on T-cell dependent immunoglobulin responses and isotype switching. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Maliszewski, J. Immunol. 144:3028-3033, 1990; Mond, J.J. and Brunswick, M Assays for B Cell Function: *In vitro* Antibody Production, Vol 1 pp. 3.8.1-3.8.16 in Current Protocols in Immunology. J.E. Coligan et al Eds., John Wiley and Sons, Toronto. 1994.

The proteins encoded by the cDNAs may also be evaluated for their effect on immune effector cells, including their effect on Th1 cells and cytotoxic lymphocytes. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Chapter 3 (In Vitro Assays for Mouse Lymphocyte

Function 3.1-3.19) and Chapter 7 (Immunologic Studies in Humans) in Current Protocols in Immunology, J.E. Coligan et al. Eds., Greene Publishing Associates and Wiley-Interscience; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., J. Immunol. 149:3778-3783, 1992.

The proteins encoded by the cDNAs may also be evaluated for their effect on dendritic cell mediated activation
of naive T-cells. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Guery et al., J. Immunol. 134:536-544, 1995; Inaba et al., Journal of Experimental Medicine 173:549-559, 1991; Macatonia et al., Journal of Immunology 154:5071-5079, 1995; Porgador et al., Journal of Experimental Medicine 182:255-260, 1995; Nair et al., Journal of Virology 67:4062-4069, 1993; Huang et al., Science 264:961-965, 1994; Macatonia et al., Journal of Experimental Medicine 169:1255-1264,
10 1989; Bhardwaj et al., Journal of Clinical Investigation 94:797-807, 1994; and Inaba et al., Journal of Experimental Medicine 172:631-640, 1990.

The proteins encoded by the cDNAs may also be evaluated for their influence on the lifetime of lymphocytes.

Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Darzynkiewicz et al., Cytometry 13:795-808, 1992; Gorczyca et al., Leukemia 7:659-670, 1993; Gorczyca et al., Cancer Research 53:1945-1951, 1993; Itoh et al., Cell 66:233-243, 1991; Zacharchuk, Journal of Immunology 145:4037-4045, 1990; Zamai et al., Cytometry 14:891-897, 1993; Gorczyca et al., International Journal of Oncology 1:639-648, 1992.

Assays for proteins that influence early steps of T-cell commitment and development include, without limitation, those described in: Antica et al., Blood 84:111-117, 1994; Fine et al., Cellular immunology 155:111-122, 1994; Galy et al., Blood 85:2770-2778, 1995; Toki et al., Proc. Nat. Acad Sci. USA 88:7548-7551, 1991.

Those proteins which exhibit activity as immune system regulators activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of immune activity is beneficial. For example, the protein may be useful in the treatment of various immune deficiencies and disorders (including severe combined immunodeficiency (SCID)), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial or fungal infections, or may result from autoimmune disorders. More specifically, infectious diseases caused by viral, bacterial, fungal or other infection may be treatable using a protein of the present invention, including infections by HIV, hepatitis viruses, herpesviruses, mycobacteria, Leishmania spp., malaria spp. and various fungal infections such as candidiasis. Of course, in this regard, a protein of the present invention may also be useful where a boost to the immune system generally may be desirable, i.e., in the treatment of cancer.

Autoimmune disorders which may be treated using a protein of the present invention include, for example, connective tissue disease, multiple sclerosis, systemic lupus erythematosus, rheumatoid arthritis, autoimmune pulmonary inflammation, Guillain-Barre syndrome, autoimmune thyroiditis, insulin dependent diabetes mellitis,

myasthenia gravis, graft-versus-host disease and autoimmune inflammatory eye disease. Such a protein of the present invention may also to be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems. Other conditions, in which immune suppression is desired (including, for example, organ transplantation), may also be treatable using a protein of the present invention.

5

Using the proteins of the invention it may also be possible to regulate immune responses, in a number of ways. Down regulation may be in the form of inhibiting or blocking an immune response already in progress or may involve preventing the induction of an immune response. The functions of activated T-cells may be inhibited by suppressing T cell responses or by inducing specific tolerance in T cells, or both. Immunosuppression of T cell responses is generally an active, non-antigen-specific, process which requires continuous exposure of the T cells to the suppressive agent. 10 Tolerance, which involves inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigen-specific and persists after exposure to the tolerizing agent has ceased. Operationally, tolerance can be demonstrated by the lack of a T cell response upon reexposure to specific antigen in the absence of the tolerizing agent.

Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte 15 antigen functions (such as, for example, B7)), e.g., preventing high level lymphokine synthesis by activated T cells, will be useful in situations of tissue, skin and organ transplantation and in graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in tissue transplantation. Typically, in tissue transplants, rejection of the transplant is initiated through its recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks interaction of a B7 20 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2 activity alone or in conjunction with a monomeric form of a peptide having an activity of another B lymphocyte antigen (e.g., B7-1, B7-3) or blocking antibody), prior to transplantation can lead to the binding of the molecule to the natural ligand(s) on the immune cells without transmitting the corresponding costimulatory signal. Blocking B lymphocyte antigen function in this matter prevents cytokine synthesis by immune cells, such as T cells, and thus acts as an 25 immunosuppressant. Moreover, the lack of costimulation may also be sufficient to anergize the T cells, thereby inducing tolerance in a subject. Induction of long-term tolerance by B lymphocyte antigen-blocking reagents may avoid the necessity of repeated administration of these blocking reagents. To achieve sufficient immunosuppression or tolerance in a subject, it may also be necessary to block the function of a combination of B lymphocyte antigens.

The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be assessed 30 using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have been used to examine the immunosuppressive effects of CTLA4lg fusion proteins in vivo as described in Lenschow et al., Science 257:789-792 (1992) and Turka et al., Proc. Natl. Acad. Sci USA, 89:11102-11105 (1992). In addition, murine models

of GVHD (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function in vivo on the development of that disease.

Blocking antigen function may also be therapeutically useful for treating autoimmune diseases. Many autoimmune disorders are the result of inappropriate activation of T cells that are reactive against self tissue and which 5 promote the production of cytokines and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents which block costimulation of T cells by disrupting receptor ligand interactions of B lymphocyte antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which may be involved in the disease process. Additionally, blocking reagents may induce antigen-specific tolerance of autoreactive T cells which could lead 10 to long-term relief from the disease. The efficacy of blocking reagents in preventing or alleviating autoimmune disorders can be determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine experimental autoimmune encephalitis, systemic lupus erythmatosis in MRL/pr/pr mice or NZB hybrid mice, murine autoimmuno collagen arthritis, diabetes mellitus in OD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy. Upregulation of immune responses may be in the form of enhancing an existing immune response or eliciting an initial immune response. For example, enhancing an immune response through stimulating B lymphocyte antigen function may be useful in cases of viral infection. In addition, systemic viral diseases such as influenza, the common cold, and encephalitis might be alleviated by the administration of stimulatory 20 form of B lymphocyte antigens systemically.

Alternatively, anti-viral immune responses may be enhanced in an infected patient by removing T cells from the patient, costimulating the T cells in vitro with viral antigen-pulsed APCs either expressing a peptide of the present invention or together with a stimulatory form of a soluble peptide of the present invention and reintroducing the in vitro activated T cells into the patient. The infected cells would now be capable of delivering a costimulatory signal to T cells in vivo, thereby activating the T cells.

In another application, up regulation or enhancement of antigen function (preferably B lymphocyte antigen function) may be useful in the induction of tumor immunity. Tumor cells (e.g., sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with a nucleic acid encoding at least one peptide of the present invention can be administered to a subject to overcome tumor-specific tolerance in the subject. If desired, the tumor cell can be 30 transfected to express a combination of peptides. For example, tumor cells obtained from a patient can be transfected ex vivo with an expression vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor cells are returned to the patient to result in expression of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection in vivo.

The presence of the peptide of the present invention having the activity of a B lymphocyte antigen(s) on the surface of the tumor cell provides the necessary costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack MHC class I or MHC class II molecules, or which fail to reexpress sufficient amounts of MHC class I or MHC class II molecules, can be transfected with nucleic acids encoding all or a portion of (e.g., a cytoplasmic-domain truncated portion) of an MHC class I α chain protein and β₂ macroglobulin protein or an MHC class II α chain protein and an MHC class II β chain protein to thereby express MHC class I or MHC class II proteins on the cell surface. Expression of the appropriate class II or class II MHC in conjunction with a peptide having the activity of a B lymphocyte antigen (e.g., B7·1, B7·2, B7·3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a gene encoding an antisense construct which blocks expression of an MHC class II associated protein, such as the invariant chain, can also be cotransfected with a DNA encoding a peptide having the activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a T cell mediated immune response in a human subject may be sufficient to overcome tumor-specific tolerance in the subject. Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

EXAMPLE 34

Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Hematopoiesis Regulating Activity

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their hematopoiesis regulating activity. For example, the effect of the proteins on embryonic stem cell differentiation may be evaluated.

Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Johansson et al. Cellular Biology 15:141-151, 1995; Keller et al., Molecular and Cellular Biology 13:473-486, 1993; McClanahan et al., Blood 81:2903-2915, 1993.

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their influence on the lifetime of stem cells and stem cell differentiation. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Freshney, M.G. Methylcellulose Colony Forming Assays, in Culture of Hematopoietic Cells. R.I. Freshney, et al. Eds. pp. 265-268, Wiley-Liss, Inc., New York, NY. 1994; Hirayama et al., Proc. Natl. Acad. Sci. USA 89:5907-5911, 1992; McNiece, I.K. and Briddell, R.A. Primitive Hematopoietic Colony Forming Cells with High Proliferative Potential, in Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 23-39, Wiley-Liss, Inc., New York, NY. 1994; Neben et al., Experimental Hematology 22:353-359, 1994; Ploemacher, R.E. Cobblestone Area Forming Cell Assay, In Culture of Hematopoietic Cells. R.I. Freshney, et al. Eds. pp. 1-21, Wiley-Liss, Inc., New York, NY. 1994; Spooncer, E., Dexter, M. and Allen, T. Long Term Bone Marrow Cultures in the Presence of Stromal Cells, in Culture of Hematopoietic Cells. R.I. Freshney, et al. Eds.

pp. 163-179, Wiley-Liss, Inc., New York, NY. 1994; and Sutherland, H.J. Long Term Culture Initiating Cell Assay, in Culture of Hematopoietic Cells. R.I. Freshney, et al. Eds. pp. 139-162, Wiley-Liss, Inc., New York, NY. 1994.

Those proteins which exhibit hematopoiesis regulatory activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of hematopoeisis is beneficial. For example, a protein of the present 5 invention may be useful in regulation of hematopoiesis and, consequently, in the treatment of myeloid or lymphoid cell deficiencies. Even marginal biological activity in support of colony forming cells or of factor-dependent cell lines indicates involvement in regulating hematopoiesis, e.g. in supporting the growth and proliferation of erythroid progenitor cells alone or in combination with other cytokines, thereby indicating utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or erythroid 10 cells; in supporting the growth and proliferation of myeloid cells such as granulocytes and monocytes/macrophages (i.e., traditional CSF activity) useful, for example, in conjunction with chemotherapy to prevent or treat consequent myelosuppression; in supporting the growth and proliferation of megakaryocytes and consequently of platelets thereby allowing prevention or treatment of various platelet disorders such as thrombocytopenia, and generally for use in place of or complimentary to platelet transfusions; and/or in supporting the growth and proliferation of hematopoietic stem 15 cells which are capable of maturing to any and all of the above-mentioned hematopoietic cells and therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantion, including, without limitation, aplastic anemia and paroxysmal nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either in-vivo or ex-vivo (i.e., in conjunction with bone marrow transplantation or with peripheral progenitor cell transplantation (homologous or heterologous)) as normal cells or 20 genetically manipulated for gene therapy. Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

EXAMPLE 35

Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Tissue Growth

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their effect on tissue growth. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in International Patent Publication No. W095/16035, International Patent Publication No. W095/05846 and International Patent Publication No. W091/07491.

Assays for wound healing activity include, without limitation, those described in: Winter, <u>Epidermal Wound</u>

30 <u>Healing</u>, pps. 71-112 (Maibach, H1 and Rovee, DT, eds.), Year Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, J. Invest. Dermatol 71:382-84 (1978).

Those proteins which are involved in the regulation of tissue growth may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of tissue growth is beneficial. For example, a protein of the present invention also may have utility in compositions used for bone, cartilage, tendon, ligament and/or

WO 99/31236 PCT/IB98/02122

-59-

nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.

A protein of the present invention, which induces cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and 5 other animals. Such a preparation employing a protein of the invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. De novo bone formation induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein of this invention may also be used in the treatment of periodontal disease, and in other tooth repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of progenitors of bone-forming cells. A protein of the invention may also be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

15

30

Another category of tissue regeneration activity that may be attributable to the protein of the present invention is tendon/ligament formation. A protein of the present invention, which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a tendon/ligament-like tissue inducing protein may have prophylactic use in preventing damage to 20 tendon or ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue formation induced by a composition of the present invention contributes to the repair of congenital, trauma induced, or other tendon or ligament defects of other origin, and is also useful in cosmetic plastic surgery for attachment or repair of tendons or ligaments. The compositions of the present invention may provide an environment to attract tendon- or ligament-forming cells, stimulate 25 growth of tendon- or ligament-forming cells, induce differentiation of progenitors of tendon- or ligament-forming cells, or induce growth of tendon/ligament cells or progenitors ex vivo for return in vivo to effect tissue repair. The compositions of the invention may also be useful in the treatment of tendinitis, carpal tunnel syndrome and other tendon or ligament defects. The compositions may also include an appropriate matrix and/or sequestering agent as a carrier as is well known in the art.

The protein of the present invention may also be useful for proliferation of neural cells and for regeneration of nerve and brain tissue, i.e., for the treatment of central and peripheral nervous system diseases and neuropathies, as well as mechanical and traumatic disorders, which involve degeneration, death or trauma to neural cells or nerve tissue. More specifically, a protein may be used in the treatment of diseases of the peripheral nervous system, such as peripheral nerve injuries, peripheral neuropathy and localized neuropathies, and central nervous system diseases, such as Alzheimer's, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Orager syndrome. Further conditions which may be treated in accordance with the present invention include mechanical and traumatic disorders, such as spinal cord disorders, head trauma and cerebrovascular diseases such as stroke. Peripheral neuropathies resulting from chemotherapy or other medical therapies may also be treatable using a protein of the invention.

Proteins of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds, and the like.

It is expected that a protein of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium) muscle

(smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition or modulation of fibrotic scarring to allow normal tissue to generate. A protein of the invention may also exhibit angiogenic activity.

A protein of the present invention may also be useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokinc damage.

A protein of the present invention may also be useful for promoting or inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

20

15

5

EXAMPLE 36

Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Reproductive Hormones or Cell Movement

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their ability to regulate reproductive hormones, such as follicle stimulating hormone. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Vale et al., Endocrinology 91:562-572, 1972; Ling et al., Nature 321:779-782, 1986; Vale et al., Nature 321:776-779, 1986; Mason et al., Nature 318:659-663, 1985; Forage et al., Proc. Natl. Acad. Sci. USA 83:3091-3095, 1986. Chapter 6.12 (Measurement of Alpha and Beta Chemokines) Current Protocols in Immunology, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Intersciece; Taub et al. J. Clin. Invest. 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Muller et al. Eur. J. Immunol. 25:1744-1748; Gruber et al. J. of Immunol. 152:5860-5867, 1994; Johnston et al. J. of Immunol. 153:1762-1768, 1994.

Those proteins which exhibit activity as reproductive hormones or regulators of cell movement may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of reproductive hormones or cell movement are beneficial. For example, a protein of the present invention may also exhibit activin- or inhibin-related

activities. Inhibins are characterized by their ability to inhibit the release of follicle stimulating hormone (FSH), while activins are characterized by their ability to stimulate the release of folic stimulating hormone (FSH). Thus, a protein of the present invention, alone or in heterodimers with a member of the inhibin α family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in female mammals and decrease spermatogenesis in male mammals.

Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein of the invention, as a homodimer or as a heterodimer with other protein subunits of the inhibin-B group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from cells of the anterior pituitary. See, for example, United States Patent 4,798,885. A protein of the invention may also be useful for advancement of the onset of fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

EXAMPLE 36A

15 <u>Assaying the Proteins Expressed from Extended cDNAs or</u>

25

30

Portions Thereof for Chemotactic/Chemokinetic Activity

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for chemotactic/chemokinetic activity. For example, a protein of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, cosinophils, epithelial and/or endothelial cells. Chemotactic and chmokinetic proteins can be used to mobilize or attract a desired cell population to a desired site of action. Chemotactic or chemokinetic proteins provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized infections. For example, attraction of lymphocytes, monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or peptide has chemotactic activity for a particular cell population if it can stimulate, directly or indirectly, the directed orientation or movement of such cell population. Preferably, the protein or peptide has the ability to directly stimulate directed movement of cells. Whether a particular protein has chemotactic activity for a population of cells can be readily determined by employing such protein or peptide in any known assay for cell chemotaxis.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins that induce or prevent chemotaxis) consist of assays that measure the ability of a protein to induce the migration of cells across a membrane as well as the ability of a protein to induce the adhension of one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those described in: Current Protocols in Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W. Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 6.12,

Measurement of alpha and beta Chemokincs 6.12.1-6.12.28; Taub et al. J. Clin. Invest. 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Mueller et al Eur. J. Immunol. 25:1744-1748; Gruber et al. J. of Immunol. 152:5860-5867, 1994; Johnston et al. J. of Immunol, 153:1762-1768, 1994.

EXAMPLE 37

5

Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Blood Clotting

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their effects on blood clotting. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Linet et al., J. Clin. Pharmacol. 26:131-140, 1986; Burdick et al., Thrombosis Res.

10 45:413-419, 1987; Humphrey et al., Fibrinolysis 5:71-79 (1991); Schaub, Prostaglandins 35:467-474, 1988.

Those proteins which are involved in the regulation of blood clotting may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of blood clotting is beneficial. For example, a protein of the invention may also exhibit hemostatic or thrombolytic activity. As a result, such a protein is expected to be useful in treatment of various coagulations disorders (including hereditary disorders, such as hemophilias) or to enhance coagulation and other hemostatic events in treating wounds resulting from trauma, surgery or other causes. A protein of the invention may also be useful for dissolving or inhibiting formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as, for example, infarction of cardiac and central nervous system vessels (e.g., stroke). Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

EXAMPLE 38

Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Involvement in Receptor/Ligand Interactions

The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for their involvement in receptor/ligand interactions. Numerous assays for such involvement are familiar to those skilled in the art, including the assays disclosed in the following references: Chapter 7.28 (Measurement of Cellular Adhesion under Static Conditions 7.28.1-7.28.22) in Current Protocols in Immunology, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Interscience; Takai et al., Proc. Natl. Acad. Sci. USA 84:6864-6868, 1987; Bierer et al., J. Exp. Med. 168:1145-1156, 1988; Rosenstein et al., J. Exp. Med. 169:149-160, 1989; Stoltenborg et al., J. Immunol. Methods 175:59-68, 1994; Stitt et al., Cell 80:661-670, 1995; Gyuris et al., Cell 75:791-803, 1993.

For example, the proteins of the present invention may also demonstrate activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands, receptors involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion

molecules (such as selectins, integrins and their ligands) and receptor/ligand pairs involved in antigen presentation, antigen recognition and development of cellular and humoral immune respones). Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction. A protein of the present invention (including, without limitation, fragments of receptors and ligands) may themselves be useful as inhibitors of receptor/ligand interactions.

EXAMPLE 38A

Assaying the Proteins Expressed from Extended cDNAs or Portions

Thereof for Anti-Inflammatory Activity

The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for anti-inflammatory activity. The anti-inflammatory activity may be achieved by providing a stimulus to cells involved in the inflammatory response, by inhibiting or promoting cell-cell interactions (such as, for example, cell adhesion), by inhibiting or promoting chemotaxis of cells involved in the inflammatory process, inhibiting or promoting cell extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins exhibiting such activities can be used to treat inflammatory conditions including chronic or acute conditions), including without limitation inflammation associated with infection (such as septic shock, sepsis or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusioninury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine or chemokine-induced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of cytokines such as TNF or IL-1. Proteins of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material.

20

30

EXAMPLE 38B

Assaying the Proteins Expressed from Extended cDNAs or

Portions Thereof for Tumor Inhibition Activity

The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for tumor inhibition activity. In addition to the activities described above for immunological treatment or prevention of tumors, a protein of the invention may exhibit other anti-tumor activities. A protein may inhibit tumor growth directly or indirectly (such as, for example, via ADCC). A protein may exhibit its tumor inhibitory activity by acting on tumor tissue or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other factors, agents or cell types which inhibit tumor growth, or by suppressing, climinating or inhibiting factors, agents or cell types which promote tumor growth.

A protein of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body part size or shape (such as, for example, breast augmentation or diminution, change in bone form or shape); effecting biorhythms or

circadian cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or climination of dietary fat, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent behaviors; providing analgesic effects or other pain reducing effects; promoting differentiation and growth of embryonic stem cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis); immunoglobulin-like activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antigen in a vaccine composition to raise an immune response against such protein or another material or entity which is cross-reactive with such protein.

EXAMPLE 39

Identification of Proteins which Interact with Polypeptides Encoded by Extended cDNAs

Proteins which interact with the polypeptides encoded by extended cDNAs or portions thereof, such as

receptor proteins, may be identified using two hybrid systems such as the Matchmaker Two Hybrid System 2 (Catalog No. K1604-1, Clontech). As described in the manual accompanying the Matchmaker Two Hybrid System 2 (Catalog No. K1604-1, Clontech), the extended cDNAs or portions thereof, are inserted into an expression vector such that they are in frame with DNA encoding the DNA binding domain of the yeast transcriptional activator GAL4. cDNAs in a cDNA library which encode proteins which might interact with the polypeptides encoded by the extended cDNAs or portions thereof

are inserted into a second expression vector such that they are in frame with DNA encoding the activation domain of GAL4. The two expression plasmids are transformed into yeast and the yeast are plated on selection medium which selects for expression of selectable markers on each of the expression vectors as well as GAL4 dependent expression of the HIS3 gene. Transformants capable of growing on medium lacking histidine are screened for GAL4 dependent lacZ expression. Those cells which are positive in both the histidine selection and the lacZ assay contain plasmids encoding proteins which interact with the polypeptide encoded by the extended cDNAs or portions thereof.

Alternatively, the system described in Lustig et al., Methods in Enzymology 283: 83-99 (1997) may be used for identifying molecules which interact with the polypeptides encoded by extended cDNAs. In such systems, *in vitro* transcription reactions are performed on a pool of vectors containing extended cDNA inserts cloned downstream of a promoter which drives *in vitro* transcription. The resulting pools of mRNAs are introduced into *Xenopus laevis* oocytes.

30 The oocytes are then assayed for a desired activity.

Alternatively, the pooled *in vitro* transcription products produced as described above may be translated *in vitro*.

The pooled *in vitro* translation products can be assayed for a desired activity or for interaction with a known polypeptide.

Proteins or other molecules interacting with polypeptides encoded by extended cDNAs can be found by a variety of additional techniques. In one method, affinity columns containing the polypeptide encoded by the extended cDNA or a portion thereof can be constructed. In some versions, of this method the affinity column contains chimeric proteins in which the protein encoded by the extended cDNA or a portion thereof is fused to glutathione S-transferase.

5 A mixture of cellular proteins or pool of expressed proteins as described above and is applied to the affinity column. Proteins interacting with the polypeptide attached to the column can then be isolated and analyzed on 2-D electrophoresis gel as described in Ramunsen et al. Electrophoresis, 18, 588-598 (1997). Alternatively, the proteins retained on the affinity column can be purified by electrophoresis based methods and sequenced. The same method can be used to isolate antibodies, to screen phage display products, or to screen phage display human antibodies.

Proteins interacting with polypeptides encoded by extended cDNAs or portions thereof can also be screened by using an Optical Biosensor as described in Edwards & Leatherbarrow, Analytical Biochemistry, 246, 1-6 (1997). The main advantage of the method is that it allows the determination of the association rate between the protein and other interacting molecules. Thus, it is possible to specifically select interacting molecules with a high or low association rate. Typically a target molecule is linked to the sensor surface (through a carboxymethi dextran matrix) and a sample of test molecules is placed in contact with the target molecules. The binding of a test molecule to the target molecule causes a change in the refractive index and/ or thickness. This change is detected by the Biosensor provided it occurs in the evanescent field (which extend a few hundred manometers from the sensor surface). In these screening assays, the target molecule can be one of the polypeptides encoded by extended cDNAs or a portion thereof and the test sample can be a collection of proteins extracted from tissues or cells, a pool of expressed proteins, combinatorial peptide and/ or chemical libraries, or phage displayed peptides. The tissues or cells from which the test proteins are extracted can originate from any species.

In other methods, a target protein is immobilized and the test population is a collection of unique polypeptides encoded by the extended cDNAs or portions thereof.

To study the interaction of the proteins encoded by the extended cDNAs or portions thereof with drugs, the microdialysis coupled to HPLC method described by Wang et al., Chromatographia, 44, 205-208(1997) or the affinity capillary electrophoresis method described by Busch et al., J. Chromatogr. 777:311-328 (1997), the disclosures of which are incorporated herein by referenc can be used.

The system described in U.S. Patent No. 5,654,150 may also be used to identify molecules which interact with the polypeptides encoded by the extended cDNAs. In this system, pools of extended cDNAs are transcribed and translated *in vitro* and the reaction products are assayed for interaction with a known polypeptide or antibody.

It will be appreciated by those skilled in the art that the proteins expressed from the extended cDNAs or portions may be assayed for numerous activities in addition to those specifically enumerated above. For example, the expressed proteins may be evaluated for applications involving control and regulation of inflammation, tumor

proliferation or metastasis, infection, or other clinical conditions. In addition, the proteins expressed from the extended cDNAs or portions thereof may be useful as nutritional agents or cosmetic agents.

The proteins expressed from the extended cDNAs or portions thereof may be used to generate antibodies capable of specifically binding to the expressed protein or fragments thereof as described in Example 40 below. The antibodies may capable of binding a full length protein encoded by one of the sequences of SEQ ID NOs. 40-140 and 242-377, a mature protein encoded by one of the sequences of SEQ ID NOs. 40-140 and 242-377, or a signal peptide encoded by one of the sequences of SEQ ID Nos. 40-140 and 242-377. Alternatively, the antibodies may be capable of binding fragments of the proteins expressed from the extended cDNAs which comprise at least 10 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513. In some embodiments, the antibodies may be capable of binding fragments of the proteins expressed from the extended cDNAs which comprise at least 15 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513. In other embodiments, the antibodies may be capable of binding fragments of the proteins expressed from the extended cDNAs which comprise at least 25 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513. In further embodiments, the antibodies may be capable of binding fragments of the proteins expressed from the extended cDNAs which comprise at least 40 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513.

EXAMPLE 40

Production of an Antibody to a Human Protein

Substantially pure protein or polypeptide is isolated from the transfected or transformed cells as described in Example 30. The concentration of protein in the final preparation is adjusted, for example, by concentration on an Amicon filter device, to the level of a few micrograms/ml. Monoclonal or polyclonal antibody to the protein can then be prepared as follows:

A. Monoclonal Antibody Production by Hybridoma Fusion

Monoclonal antibody to epitopes of any of the peptides identified and isolated as described can be prepared from murine hybridomas according to the classical method of Kohler, G. and Milstein, C., Nature 256:495 (1975) or derivative methods thereof. Briefly, a mouse is repetitively inoculated with a few micrograms of the selected protein or peptides derived therefrom over a period of a few weeks. The mouse is then sacrificed, and the antibody producing cells of the spleen isolated. The spleen cells are fused by means of polyethylene glycol with mouse myeloma cells, and the excess unfused cells destroyed by growth of the system on selective media comprising aminopterin (HAT media). The successfully fused cells are diluted and aliquots of the dilution placed in wells of a microtiter plate where growth of the culture is continued. Antibody-producing clones are identified by detection of antibody in the supernatant fluid of the wells by immunoassay procedures, such as Elisa, as originally described by Engvall, E., Meth. Enzymol. 70:419 (1980), and derivative methods thereof. Selected positive clones can be expanded and their monoclonal antibody product harvested for use. Detailed procedures for monoclonal antibody production are described in Davis, L. et al. Besic Methods in Molecular Biology Elsevier, New York. Section 21-2.

B. Polyclonal Antibody Production by Immunization

Polyclonal antiserum containing antibodies to heterogenous epitopes of a single protein can be prepared by immunizing suitable animals with the expressed protein or peptides derived therefrom described above, which can be unmodified or modified to enhance immunogenicity. Effective polyclonal antibody production is affected by many factors 5 related both to the antigen and the host species. For example, small molecules tend to be less immunogenic than others and may require the use of carriers and adjuvant. Also, host animals vary in response to site of inoculations and dose, with both inadequate or excessive doses of antigen resulting in low titer antisera. Small doses (ng level) of antigen administered at multiple intradermal sites appears to be most reliable. An effective immunization protocol for rabbits can be found in Vaitukaitis, J. et al. J. Clin. Endocrinol. Metab. 33:988-991 (1971).

Booster injections can be given at regular intervals, and antiserum harvested when antibody titer thereof, as determined semi-quantitatively, for example, by double immunodiffusion in agar against known concentrations of the antigen, begins to fall. See, for example, Ouchterlony, O. et al., Chap. 19 in: Handbook of Experimental Immunology D. Wier (ed) Blackwell (1973). Plateau concentration of antibody is usually in the range of 0.1 to 0.2 mg/ml of serum (about 12 µM). Affinity of the antisera for the antigen is determined by preparing competitive binding curves, as 15 described, for example, by Fisher, D., Chap. 42 in: Manual of Clinical Immunology, 2d Ed. (Rose and Friedman, Eds.) Amer. Soc. For Microbiol., Washington, D.C. (1980).

Antibody preparations prepared according to either protocol are useful in quantitative immunoassays which determine concentrations of antigen-bearing substances in biological samples; they are also used semi-quantitatively or qualitatively to identify the presence of antigen in a biological sample. The antibodies may also be used in therapeutic 20 compositions for killing cells expressing the protein or reducing the levels of the protein in the body.

V. Use of Extended cDNAs or Portions Thereof as Reagents

The extended cDNAs of the present invention may be used as reagents in isolation procedures, diagnostic assays, and forensic procedures. For example, sequences from the extended cDNAs (or genomic DNAs obtainable 25 therefrom) may be detectably labeled and used as probes to isolate other sequences capable of hybridizing to them. In addition, sequences from the extended cDNAs (or genomic DNAs obtainable therefrom) may be used to design PCR primers to be used in isolation, diagnostic, or forensic procedures.

EXAMPLE 41

Preparation of PCR Primers and Amplification of DNA

30 The extended cDNAs (or genomic DNAs obtainable therefrom) may be used to prepare PCR primers for a variety of applications, including isolation procedures for cloning nucleic acids capable of hybridizing to such sequences, diagnostic techniques and forensic techniques. The PCR primers are at least 10 bases, and preferably at least 12, 15, or 17 bases in length. More preferably, the PCR primers are at least 20-30 bases in length. In some embodiments, the PCR primers may be more than 30 bases in length. It is preferred that the primer pairs have approximately the same G/C

ratio, so that melting temperatures are approximately the same. A variety of PCR techniques are familiar to those skilled in the art. For a review of PCR technology, see Molecular Cloning to Genetic Engineering White, B.A. Ed. in Methods in Molecular Biology 67: Humana Press, Totowa 1997. In each of these PCR procedures, PCR primers on either side of the nucleic acid sequences to be amplified are added to a suitably prepared nucleic acid sample along with dNTPs and a thermostable polymerase such as Taq polymerase, Pfu polymerase, or Vent polymerase. The nucleic acid in the sample is denatured and the PCR primers are specifically hybridized to complementary nucleic acid sequences in the sample. The hybridized primers are extended. Thereafter, another cycle of denaturation, hybridization, and extension is initiated. The cycles are repeated multiple times to produce an amplified fragment containing the nucleic acid sequence between the primer sites.

10

EXAMPLE 42

Use of Extended cDNAs as Probes

Probes derived from extended cDNAs or portions thereof (or genomic DNAs obtainable therefrom) may be labeled with detectable labels familiar to those skilled in the art, including radioisotopes and non-radioactive labels, to provide a detectable probe. The detectable probe may be single stranded or double stranded and may be made using techniques known in the art, including in vitro transcription, nick translation, or kinase reactions. A nucleic acid sample containing a sequence capable of hybridizing to the labeled probe is contacted with the labeled probe. If the nucleic acid in the sample is double stranded, it may be denatured prior to contacting the probe. In some applications, the nucleic acid sample may be immobilized on a surface such as a nitrocellulose or nylon membrane. The nucleic acid sample may comprise nucleic acids obtained from a variety of sources, including genomic DNA, cDNA libraries, RNA, or tissue samples.

Procedures used to detect the presence of nucleic acids capable of hybridizing to the detectable probe include well known techniques such as Southern blotting, Northern blotting, dot blotting, colony hybridization, and plaque hybridization. In some applications, the nucleic acid capable of hybridizing to the labeled probe may be cloned into vectors such as expression vectors, sequencing vectors, or in vitro transcription vectors to facilitate the characterization and expression of the hybridizing nucleic acids in the sample. For example, such techniques may be used to isolate and clone sequences in a genomic library or cDNA library which are capable of hybridizing to the detectable probe as described in Example 30 above.

PCR primers made as described in Example 41 above may be used in forensic analyses, such as the DNA fingerprinting techniques described in Examples 43-47 below. Such analyses may utilize detectable probes or primers based on the sequences of the extended cDNAs isolated using the 5' ESTs (or genomic DNAs obtainable therefrom).

EXAMPLE 43

Forensic Matching by DNA Sequencing

In one exemplary method, DNA samples are isolated from forensic specimens of, for example, hair, semen, blood or skin cells by conventional methods. A panel of PCR primers based on a number of the extended cDNAs (or

genomic DNAs obtainable therefrom), is then utilized in accordance with Example 41 to amplify DNA of approximately 100-200 bases in length from the forensic specimen. Corresponding sequences are obtained from a test subject. Each of these identification DNAs is then sequenced using standard techniques, and a simple database comparison determines the differences, if any, between the sequences from the subject and those from the sample. Statistically significant differences between the suspect's DNA sequences and those from the sample conclusively prove a lack of identity. This lack of identity can be proven, for example, with only one sequence. Identity, on the other hand, should be demonstrated with a large number of sequences, all matching. Preferably, a minimum of 50 statistically identical sequences of 100 bases in length are used to prove identity between the suspect and the sample.

EXAMPLE 44

10

Positive Identification by DNA Sequencing

The technique outlined in the previous example may also be used on a larger scale to provide a unique fingerprint-type identification of any individual. In this technique, primers are prepared from a large number of sequences from Table IV and the appended sequence listing. Preferably, 20 to 50 different primers are used. These primers are used to obtain a corresponding number of PCR-generated DNA segments from the individual in question in accordance with Example 41. Each of these DNA segments is sequenced, using the methods set forth in Example 43. The database of sequences generated through this procedure uniquely identifies the individual from whom the sequences were obtained. The same panel of primers may then be used at any later time to absolutely correlate tissue or other biological specimen with that individual.

EXAMPLE 45

20

Southern Blot Forensic Identification

The procedure of Example 44 is repeated to obtain a panel of at least 10 amplified sequences from an individual and a specimen. Preferably, the panel contains at least 50 amplified sequences. More preferably, the panel contains 100 amplified sequences. In some embodiments, the panel contains 200 amplified sequences. This PCR-generated DNA is then digested with one or a combination of, preferably, four base specific restriction enzymes. Such enzymes are commercially available and known to those of skill in the art. After digestion, the resultant gene fragments are size separated in multiple duplicate wells on an agarose gel and transferred to nitrocellulose using Southern blotting techniques well known to those with skill in the art. For a review of Southern blotting see Davis et al. (Basic Methods in Molecular Biology, 1986, Elsevier Press. pp 62-65).

A panel of probes based on the sequences of the extended cDNAs (or genomic DNAs obtainable therefrom), or fragments thereof of at least 10 bases, are radioactively or colorimetrically labeled using methods known in the art, such as nick translation or end labeling, and hybridized to the Southern blot using techniques known in the art (Davis et al., supra). Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). More preferably, the probe comprises at least 20-30 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In some embodiments, the probe comprises more than 30

nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In other embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom).

Preferably, at least 5 to 10 of these labeled probes are used, and more preferably at least about 20 or 30 are used to provide a unique pattern. The resultant bands appearing from the hybridization of a large sample of extended cDNAs (or genomic DNAs obtainable therefrom) will be a unique identifier. Since the restriction enzyme cleavage will be different for every individual, the band pattern on the Southern blot will also be unique. Increasing the number of extended cDNA probes will provide a statistically higher level of confidence in the identification since there will be an increased number of sets of bands used for identification.

10

25

EXAMPLE 46

Dot Blot Identification Procedure

Another technique for identifying individuals using the extended cDNA sequences disclosed herein utilizes a dot blot hybridization technique.

Genomic DNA is isolated from nuclei of subject to be identified. Oligonucleotide probes of approximately 30 bp in length are synthesized that correspond to at least 10, preferably 50 sequences from the extended cDNAs or genomic DNAs obtainable therefrom. The probes are used to hybridize to the genomic DNA through conditions known to those in the art. The oligonucleotides are end labeled with P32 using polynucleotide kinase (Pharmacia). Dot Blots are created by spotting the genomic DNA onto nitrocellulose or the like using a vacuum dot blot manifold (BioRad, Richmond California). The nitrocellulose filter containing the genomic sequences is baked or UV linked to the filter, prehybridized and hybridized with labeled probe using techniques known in the art (Davis et al. supra). The 32P labeled DNA fragments are sequentially hybridized with successively stringent conditions to detect minimal differences between the 30 bp sequence and the DNA. Tetramethylammonium chloride is useful for identifying clones containing small numbers of nucleotide mismatches (Wood et al., Proc. Natl. Acad. Sci. USA 82(6):1585-1588 (1985)). A unique pattern of dots distinguishes one individual from another individual.

Extended cDNAs or oligonucleotides containing at least 10 consecutive bases from these sequences can be used as probes in the following alternative fingerprinting technique. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). More preferably, the probe comprises at least 20-30 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In some embodiments, the probe comprises more than 30 nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In other embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom).

Preferably, a plurality of probes having sequences from different genes are used in the alternative fingerprinting technique. Example 47 below provides a representative alternative fingerprinting procedure in which the probes are derived from extended cDNAs.

EXAMPLE 47

5

Alternative "Fingerprint" Identification Technique

20-mer oligonucleotides are prepared from a large number, e.g. 50, 100, or 200, of extended cDNA sequences (or genomic DNAs obtainable therefrom) using commercially available oligonucleotide services such as Genset, Paris, France. Cell samples from the test subject are processed for DNA using techniques well known to those with skill in the art. The nucleic acid is digested with restriction enzymes such as EcoRI and Xbal. Following digestion, samples are applied to wells for electrophoresis. The procedure, as known in the art, may be modified to accommodate polyacrylamide electrophoresis, however in this example, samples containing 5 ug of DNA are loaded into wells and separated on 0.8% agarose gels. The gels are transferred onto nitrocellulose using standard Southern blotting techniques.

10 ng of each of the oligonucleotides are pooled and end-labeled with P³². The nitrocellulose is prehybridized with blocking solution and hybridized with the labeled probes. Following hybridization and washing, the nitrocellulose filter is exposed to X-Omat AR X-ray film. The resulting hybridization pattern will be unique for each individual.

It is additionally contemplated within this example that the number of probe sequences used can be varied for additional accuracy or clarity.

The antibodies generated in Examples 30 and 40 above may be used to identify the tissue type or cell species 20 from which a sample is derived as described above.

EXAMPLE 48

Identification of Tissue Types or Cell Species by Means of

Labeled Tissue Specific Antibodies

Identification of specific tissues is accomplished by the visualization of tissue specific antigens by means of
antibody preparations according to Examples 30 and 40 which are conjugated, directly or indirectly to a detectable
marker. Selected labeled antibody species bind to their specific antigen binding partner in tissue sections, cell
suspensions, or in extracts of soluble proteins from a tissue sample to provide a pattern for qualitative or semiqualitative interpretation.

Antisera for these procedures must have a potency exceeding that of the native preparation, and for that
reason, antibodies are concentrated to a mg/ml level by isolation of the gamma globulin fraction, for example, by ionexchange chromatography or by ammonium sulfate fractionation. Also, to provide the most specific antisera, unwanted
antibodies, for example to common proteins, must be removed from the gamma globulin fraction, for example by means
of insoluble immunoabsorbents, before the antibodies are labeled with the marker. Either monoclonal or heterologous
antisera is suitable for either procedure.

20

A. Immunohistochemical Techniques

Purified, high-titer antibodies, prepared as described above, are conjugated to a detectable marker, as described, for example, by Fudenberg, H., Chap. 26 in: Basic 503 Clinical Immunology, 3rd Ed. Lange, Los Altos, California (1980) or Rose, N. et al., Chap. 12 in: Methods in Immunodiagnosis, 2d Ed. John Wiley 503 Sons, New York (1980).

A fluorescent marker, either fluorescein or rhodamine, is preferred, but antibodies can also be labeled with an enzyme that supports a color producing reaction with a substrate, such as horseradish peroxidase. Markers can be added to tissue-bound antibody in a second step, as described below. Alternatively, the specific antitissue antibodies can be labeled with ferritin or other electron dense particles, and localization of the ferritin coupled antigen-antibody complexes achieved by means of an electron microscope. In yet another approach, the antibodies are radiolabeled, with, for example ¹²⁵I, and detected by overlaying the antibody treated preparation with photographic emulsion.

Preparations to carry out the procedures can comprise monoclonal or polyclonal antibodies to a single protein or peptide identified as specific to a tissue type, for example, brain tissue, or antibody preparations to several antigenically distinct tissue specific antigens can be used in panels, independently or in mixtures, as required.

Tissue sections and cell suspensions are prepared for immunohistochemical examination according to common histological techniques. Multiple cryostat sections (about 4 μ m, unfixed) of the unknown tissue and known control, are mounted and each slide covered with different dilutions of the antibody preparation. Sections of known and unknown tissues should also be treated with preparations to provide a positive control, a negative control, for example, pre-immune sera, and a control for non-specific staining, for example, buffer.

Treated sections are incubated in a humid chamber for 30 min at room temperature, rinsed, then washed in buffer for 30-45 min. Excess fluid is blotted away, and the marker developed.

If the tissue specific antibody was not labeled in the first incubation, it can be labeled at this time in a second antibody-antibody reaction, for example, by adding fluorescein- or enzyme-conjugated antibody against the immunoglobulin class of the antiserum-producing species, for example, fluorescein labeled antibody to mouse IgG. Such labeled sera are commercially available.

The antigen found in the tissues by the above procedure can be quantified by measuring the intensity of color or fluorescence on the tissue section, and calibrating that signal using appropriate standards.

B. Identification of Tissue Specific Soluble Proteins

The visualization of tissue specific proteins and identification of unknown tissues from that procedure is

carried out using the labeled antibody reagents and detection strategy as described for immunohistochemistry; however
the sample is prepared according to an electrophoretic technique to distribute the proteins extracted from the tissue in
an orderly array on the basis of molecular weight for detection.

A tissue sample is homogenized using a Virtis apparatus; cell suspensions are disrupted by Dounce homogenization or osmotic lysis, using detergents in either case as required to disrupt cell membranes, as is the practice

in the art. Insoluble cell components such as nuclei, microsomes, and membrane fragments are removed by ultracentrifugation, and the soluble protein-containing fraction concentrated if necessary and reserved for analysis.

A sample of the soluble protein solution is resolved into individual protein species by conventional SDS polyacrylamide electrophoresis as described, for example, by Davis, L. et al., Section 19-2 in: Basic Methods in 5 Molecular Biology (P. Leder, ed), Elsevier, New York (1986), using a range of amounts of polyacrylamide in a set of gels to resolve the entire molecular weight range of proteins to be detected in the sample. A size marker is run in parallel for purposes of estimating molecular weights of the constituent proteins. Sample size for analysis is a convenient volume of from 5 to55 µl, and containing from about 1 to 100 µg protein. An aliquot of each of the resolved proteins is transferred by blotting to a nitrocellulose filter paper, a process that maintains the pattern of resolution. Multiple copies 10 are prepared. The procedure, known as Western Blot Analysis, is well described in Davis, L. et al., (above) Section 19-3. One set of nitrocellulose blots is stained with Coomassie Blue dye to visualize the entire set of proteins for comparison with the antibody bound proteins. The remaining nitrocellulose filters are then incubated with a solution of one or more specific antisera to tissue specific proteins prepared as described in Examples 30 and 40. In this procedure, as in procedure A above, appropriate positive and negative sample and reagent controls are run.

In either procedure A or B, a detectable label can be attached to the primary tissue antigen-primary antibody complex according to various strategies and permutations thereof. In a straightforward approach, the primary specific antibody can be labeled; alternatively, the unlabeled complex can be bound by a labeled secondary anti-IgG antibody. In other approaches, either the primary or secondary antibody is conjugated to a biotin molecule, which can, in a subsequent step, bind an avidin conjugated marker. According to yet another strategy, enzyme labeled or radioactive 20 protein A, which has the property of binding to any IgG, is bound in a final step to either the primary or secondary antibody.

The visualization of tissue specific antigen binding at levels above those seen in control tissues to one or more tissue specific antibodies, prepared from the gene sequences identified from extended cDNA sequences, can identify tissues of unknown origin, for example, forensic samples, or differentiated tumor tissue that has metastasized to foreign 25 bodily sites.

In addition to their applications in forensics and identification, extended cDNAs (or genomic DNAs obtainable therefrom) may be mapped to their chromosomal locations. Example 49 below describes radiation hybrid (RH) mapping of human chromosomal regions using extended cDNAs. Example 50 below describes a representative procedure for mapping an extended cDNA (or a genomic DNA obtainable therefrom) to its location on a human chromosome. Example 30 51 below describes mapping of extended cDNAs (or genomic DNAs obtainable therefrom) on metaphase chromosomes by Fluorescence In Situ Hybridization (FISH).

EXAMPLE 49

Radiation hybrid mapping of Extended cDNAs to the human genome

Radiation hybrid (RH) mapping is a somatic cell genetic approach that can be used for high resolution mapping of the human genome. In this approach, cell lines containing one or more human chromosomes are lethally irradiated, breaking each chromosome into fragments whose size depends on the radiation dose. These fragments are rescued by fusion with cultured rodent cells, yielding subclones containing different portions of the human genome. This technique is described by Benham et al. (*Genomics* 4:509-517, 1989) and Cox et al., (*Science* 250:245-250, 1990). The random and independent nature of the subclones permits efficient mapping of any human genome marker. Human DNA isolated from a panel of 80-100 cell lines provides a mapping reagent for ordering extended cDNAs (or genomic DNAs obtainable therefrom). In this approach, the frequency of breakage between markers is used to measure distance, allowing construction of fine resolution maps as has been done using conventional ESTs (Schuler et al., *Science* 274:540-546, 1996).

RH mapping has been used to generate a high-resolution whole genome radiation hybrid map of human chromosome 17q22-q25.3 across the genes for growth hormone (GH) and thyr.nidine kinase (TK) (Foster et al., *Genomics* 33:185-192, 1996), the region surrounding the Gorlin syndrome gene (Obermayr et al., *Eur. J. Hum. Genet.* 4:242-245, 1996), 60 loci covering the entire short arm of chromosome 12 (Raeymaekers et al., *Genomics* 29:170-178, 1995), the region of human chromosome 22 containing the neurofibromatosis type 2 locus (Frazer et al., *Genomics* 14:574-584, 1992) and 13 loci on the long arm of chromosome 5 (Warrington et al., *Genomics* 11:701-708, 1991).

EXAMPLE 50

Mapping of Extended cDNAs to Human

Chromosomes using PCR techniques

Extended cDNAs (or genomic DNAs obtainable therefrom) may be assigned to human chromosomes using PCR based methodologies. In such approaches, oligonucleotide primer pairs are designed from the extended cDNA sequence (or the sequence of a genomic DNA obtainable therefrom) to minimize the chance of amplifying through an intron. Preferably, the oligonucleotide primers are 18-23 bp in length and are designed for PCR amplification. The creation of PCR primers from known sequences is well known to those with skill in the art. For a review of PCR technology see Erlich, H.A., PCR Technology; Principles and Applications for DNA Amplification. 1992. W.H. Freeman and Co., New York.

The primers are used in polymerase chain reactions (PCR) to amplify templates from total human genomic DNA. PCR conditions are as follows: 60 ng of genomic DNA is used as a template for PCR with 80 ng of each oligonucleotide primer, 0.6 unit of Taq polymerase, and 1 µCu of a ³²P-labeled deoxycytidine triphosphate. The PCR is performed in a microplate thermocycler (Techne) under the following conditions: 30 cycles of 94°C, 1.4 min; 55°C, 2 min; and 72°C, 2 min; with a final extension at 72°C for 10 min. The amplified products are analyzed on a 6% polyacrylamide sequencing gel and visualized by autoradiography. If the length of the resulting PCR product is identical to the distance between the ends of the primer sequences in the extended cDNA from which the primers are derived, then the PCR reaction is repeated with DNA templates from two panels of human-rodent somatic cell hybrids, BIOS

PCR is used to screen a series of somatic cell hybrid cell lines containing defined sets of human chromosomes for the presence of a given extended cDNA (or genomic DNA obtainable therefrom). DNA is isolated from the somatic hybrids and used as starting templates for PCR reactions using the primer pairs from the extended cDNAs (or genomic DNAs obtainable therefrom). Only those somatic cell hybrids with chromosomes containing the human gene corresponding to the extended cDNA (or genomic DNA obtainable therefrom) will yield an amplified fragment. The extended cDNAs (or genomic DNAs obtainable therefrom) are assigned to a chromosome by analysis of the segregation pattern of PCR products from the somatic hybrid DNA templates. The single human chromosome present in all cell hybrids that give rise to an amplified fragment is the chromosome containing that extended cDNA (or genomic DNA obtainable therefrom). For a review of techniques and analysis of results from somatic cell gene mapping experiments. (See Ledbetter et al., Genomics 6:475-481 (1990).)

Alternatively, the extended cDNAs (or genomic DNAs obtainable therefrom) may be mapped to individual chromosomes using FISH as described in Example 51 below.

EXAMPLE 51

Mapping of Extended 5' ESTs to Chromosomes

Using Fluorescence in situ Hybridization

Fluorescence in situ hybridization allows the extended cDNA (or genomic DNA obtainable therefrom) to be mapped to a particular location on a given chromosome. The chromosomes to be used for fluorescence in situ hybridization techniques may be obtained from a variety of sources including cell cultures, tissues, or whole blood.

In a preferred embodiment, chromosomal localization of an extended cDNA (or genomic DNA obtainable therefrom) is obtained by FISH as described by Cherif et al. (*Proc. Natl. Acad. Sci. U.S.A.*, 87:6639-6643, 1990).

Metaphase chromosomes are prepared from phytohemagglutinin (PHA)-stimulated blood cell donors. PHA-stimulated lymphocytes from healthy males are cultured for 72 h in RPMI-1640 medium. For synchronization, methotrexate (10 µM) is added for 17 h, followed by addition of 5-bromodeoxyuridine (5-BudR, 0.1 mM) for 6 h. Colcemid (1 µg/ml) is added for the last 15 min before harvesting the cells. Cells are collected, washed in RPMI, incubated with a hypotonic solution of KCI (75 mM) at 37°C for 15 min and fixed in three changes of methanol:acetic acid (3:1). The cell suspension is dropped onto a glass slide and air dried. The extended cDNA (or genomic DNA obtainable therefrom) is labeled with biotin-16 dUTP by nick translation according to the manufacturer's instructions (Bethesda Research

Laboratories, Bethesda, MD), purified using a Sephadex G-50 column (Pharmacia, Upssala, Sweden) and precipitated.

Just prior to hybridization, the DNA pellet is dissolved in hybridization buffer (50% formamide, 2 X SSC, 10% dextran sulfate, 1 mg/ml sonicated salmon sperm DNA, pH 7) and the probe is denatured at 70°C for 5-10 min.

Slides kept at -20°C are treated for 1 h at 37°C with RNase A (100 μ g/ml), rinsed three times in 2 X SSC and dehydrated in an ethanol series. Chromosome preparations are denatured in 70% formamide, 2 X SSC for 2 min at

70°C, then dehydrated at 4°C. The slides are treated with proteinase K (10 μg/100 ml in 20 mM Tris-HCl, 2 mM CaCl₂) at 37°C for 8 min and dehydrated. The hybridization mixture containing the probe is placed on the slide, covered with a coverslip, sealed with rubber cement and incubated overnight in a humid chamber at 37°C. After hybridization and post-hybridization washes, the biotinylated probe is detected by avidin-FITC and amplified with additional layers of biotinylated goat anti-avidin and avidin-FITC. For chromosomal localization, fluorescent R-bands are obtained as previously described (Cherif et al., *supra.*). The slides are observed under a LEICA fluorescence microscope (DMRXA). Chromosomes are counterstained with propidium iodide and the fluorescent signal of the probe appears as two symmetrical yellow-green spots on both chromatids of the fluorescent R-band chromosome (red). Thus, a particular extended cDNA (or genomic DNA obtainable therefrom) may be localized to a particular cytogenetic R-band on a given thromosome.

Once the extended cDNAs (or genomic DNAs obtainable therefrom) have been assigned to particular chromosomes using the techniques described in Examples 49-51 above, they may be utilized to construct a high resolution map of the chromosomes on which they are located or to identify the chromosomes in a sample.

EXAMPLE 52

Use of Extended cDNAs to Construct or Expand Chromosome Maps

Chromosome mapping involves assigning a given unique sequence to a particular chromosome as described above. Once the unique sequence has been mapped to a given chromosome, it is ordered relative to other unique sequences located on the same chromosome. One approach to chromosome mapping utilizes a series of yeast artificial chromosomes (YACs) bearing several thousand long inserts derived from the chromosomes of the organism from which the extended cDNAs (or genomic DNAs obtainable therefrom) are obtained. This approach is described in Ramaiah Nagaraja et al. Genome Research 7:210-222, March 1997. Briefly, in this approach each chromosome is broken into overlapping pieces which are inserted into the YAC vector. The YAC inserts are screened using PCR or other methods to determine whether they include the extended cDNA (or genomic DNA obtainable therefrom) whose position is to be determined. Once an insert has been found which includes the extended cDNA (or genomic DNA obtainable therefrom), the insert can be analyzed by PCR or other methods to determine whether the insert also contains other sequences known to be on the chromosome or in the region from which the extended cDNA (or genomic DNA obtainable therefrom) was derived. This process can be repeated for each insert in the YAC library to determine the location of each of the extended cDNAs (or genomic DNAs obtainable therefrom) relative to one another and to other known chromosomal markers. In this way, a high resolution map of the distribution of numerous unique markers along each of the organisms 30 chromosomes may be obtained.

As described in Example 53 below extended cDNAs (or genomic DNAs obtainable therefrom) may also be used to identify genes associated with a particular phenotype, such as hereditary disease or drug response.

EXAMPLE 53

Identification of genes associated with hereditary diseases or drug response

20

25

30

This example illustrates an approach useful for the association of extended cDNAs (or genomic DNAs obtainable therefrom) with particular phenotypic characteristics. In this example, a particular extended cDNA (or genomic DNA obtainable therefrom) is used as a test probe to associate that extended cDNA (or genomic DNA obtainable therefrom) with a particular phenotypic characteristic.

Extended cDNAs (or genomic DNAs obtainable therefrom) are mapped to a particular location on a human chromosome using techniques such as those described in Examples 49 and 50 or other techniques known in the art. A search of Mendelian Inheritance in Man (V. McKusick, Mendelian Inheritance in Man (available on line through Johns Hopkins University Welch Medical Library) reveals the region of the human chromosome which contains the extended cDNA (or genomic DNA obtainable therefrom) to be a very gene rich region containing several known genes and several 10 diseases or phenotypes for which genes have not been identified. The gene corresponding to this extended cDNA (or genomic DNA obtainable therefrom) thus becomes an immediate candidate for each of these genetic diseases.

Cells from patients with these diseases or phenotypes are isolated and expanded in culture. PCR primers from the extended cDNA (or genomic DNA obtainable therefrom) are used to screen genomic DNA, mRNA or cDNA obtained from the patients. Extended cDNAs (or genomic DNAs obtainable therefrom) that are not amplified in the patients can 15 be positively associated with a particular disease by further analysis. Alternatively, the PCR analysis may yield fragments of different lengths when the samples are derived from an individual having the phenotype associated with the disease than when the sample is derived from a healthy individual, indicating that the gene containing the extended cDNA may be responsible for the genetic disease.

VI. Use of Extended cDNAs (or genomic DNAs obtainable therefrom) to Construct Vectors

The present extended cDNAs (or genomic DNAs obtainable therefrom) may also be used to construct secretion vectors capable of directing the secretion of the proteins encoded by genes inserted in the vectors. Such secretion vectors may facilitate the purification or enrichment of the proteins encoded by genes inserted therein by reducing the number of background proteins from which the desired protein must be purified or enriched. Exemplary secretion vectors are described in Example 54 below.

EXAMPLE 54

Construction of Secretion Vectors

The secretion vectors of the present invention include a promoter capable of directing gene expression in the host cell, tissue, or organism of interest. Such promoters include the Rous Sarcoma Virus promoter, the SV40 promoter, the human cytomegalovirus promoter, and other promoters familiar to those skilled in the art.

A signal sequence from an extended cDNA (or genomic DNA obtainable therefrom), such as one of the signal sequences in SEQ ID NOs: 40-140 and 242-377 as defined in Table IV above, is operably linked to the promoter such that the mRNA transcribed from the promoter will direct the translation of the signal peptide. The host cell, tissue, or organism may be any cell, tissue, or organism which recognizes the signal peptide encoded by the signal sequence in the

extended cDNA (or genomic DNA obtainable therefrom). Suitable hosts include mammalian cells, tissues or organisms, avian cells, tissues, or organisms, insect cells, tissues or organisms, or yeast.

In addition, the secretion vector contains cloning sites for inserting genes encoding the proteins which are to be secreted. The cloning sites facilitate the cloning of the insert gene in frame with the signal sequence such that a fusion 5 protein in which the signal peptide is fused to the protein encoded by the inserted gene is expressed from the mRNA transcribed from the promoter. The signal peptide directs the extracellular secretion of the fusion protein.

The secretion vector may be DNA or RNA and may integrate into the chromosome of the host, be stably maintained as an extrachromosomal replicon in the host, be an artificial chromosome, or be transiently present in the host. Many nucleic acid backbones suitable for use as secretion vectors are known to those skilled in the art, including 10 retroviral vectors, SV40 vectors, Bovine Papilloma Virus vectors, yeast integrating plasmids, yeast episomal plasmids, veast artificial chromosomes, human artificial chromosomes, P element vectors, baculovirus vectors, or bacterial plasmids capable of being transiently introduced into the host.

The secretion vector may also contain a polyA signal such that the polyA signal is located downstream of the gene inserted into the secretion vector.

After the gene encoding the protein for which secretion is desired is inserted into the secretion vector, the secretion vector is introduced into the host cell, tissue, or organism using calcium phosphate precipitation, DEAE-Dextran, electroporation, liposome-mediated transfection, viral particles or as naked DNA. The protein encoded by the inserted gene is then purified or enriched from the supernatant using conventional techniques such as ammonium sulfate precipitation, immunoprecipitation, immunochromatography, size exclusion chromatography, ion exchange 20 chromatography, and hplc. Alternatively, the secreted protein may be in a sufficiently enriched or pure state in the supernatant or growth media of the host to permit it to be used for its intended purpose without further enrichment.

The signal sequences may also be inserted into vectors designed for gene therapy. In such vectors, the signal sequence is operably linked to a promoter such that mRNA transcribed from the promoter encodes the signal peptide. A cloning site is located downstream of the signal sequence such that a gene encoding a protein whose secretion is 25 desired may readily be inserted into the vector and fused to the signal sequence. The vector is introduced into an appropriate host cell. The protein expressed from the promoter is secreted extracellularly, thereby producing a therapeutic effect.

The extended cDNAs or 5' ESTs may also be used to clone sequences located upstream of the extended cDNAs or 5' ESTs which are capable of regulating gene expression, including promoter sequences, enhancer sequences, and 30 other upstream sequences which influence transcription or translation levels. Once identified and cloned, these upstream regulatory sequences may be used in expression vectors designed to direct the expression of an inserted gene in a desired spatial, temporal, developmental, or quantitative fashion. Example 55 describes a method for cloning sequences upstream of the extended cDNAs or 5' ESTs.

Use of Extended cDNAs or 5' ESTs to Clone Upstream

Sequences from Genomic DNA

Sequences derived from extended cDNAs or 5' ESTs may be used to isolate the promoters of the corresponding genes using chromosome walking techniques. In one chromosome walking technique, which utilizes the 5 GenomeWalker™ kit available from Clontech, five complete genomic DNA samples are each digested with a different restriction enzyme which has a 6 base recognition site and leaves a blunt end. Following digestion, oligonucleotide adapters are ligated to each end of the resulting genomic DNA fragments.

For each of the five genomic DNA libraries, a first PCR reaction is performed according to the manufacturer's instructions using an outer adaptor primer provided in the kit and an outer gene specific primer. The gene specific primer 10 should be selected to be specific for the extended cDNA or 5' EST of interest and should have a melting temperature, length, and location in the extended cDNA or 'EST which is consistent with its use in PCR reactions. Each first PCR reaction contains 5ng of genomic DNA, 5 μ l of 10X Tth reaction buffer, 0.2 mM of each dNTP, 0.2 μ M each of outer adaptor primer and outer gene specific primer, 1.1 mM of Mg(OAc)₂, and 1 μ l of the Tth polymerase 50X mix in a total volume of 50 µl. The reaction cycle for the first PCR reaction is as follows: 1 min @ 94°C / 2 sec @ 94°C, 3 min @ 15 72°C (7 cycles) / 2 sec @ 94°C, 3 min @ 67°C (32 cycles) / 5 min @ 67°C.

The product of the first PCR reaction is diluted and used as a template for a second PCR reaction according to the manufacturer's instructions using a pair of nested primers which are located internally on the amplicon resulting from the first PCR reaction. For example, 5 μ l of the reaction product of the first PCR reaction mixture may be diluted 180 times. Reactions are made in a 50 μ l volume having a composition identical to that of the first PCR reaction except 20 the nested primers are used. The first nested primer is specific for the adaptor, and is provided with the GenomeWalker™ kit. The second nested primer is specific for the particular extended cDNA or 5' EST for which the promoter is to be cloned and should have a melting temperature, length, and location in the extended cDNA or 5' EST which is consistent with its use in PCR reactions. The reaction parameters of the second PCR reaction are as follows: 1 min @ 94°C / 2 sec @ 94°C, 3 min @ 72°C (6 cycles) / 2 sec @ 94°C, 3 min @ 67°C (25 cycles) / 5 min @ 67°C

The product of the second PCR reaction is purified, cloned, and sequenced using standard techniques. Alternatively, two or more human genomic DNA libraries can be constructed by using two or more restriction enzymes. The digested genomic DNA is cloned into vectors which can be converted into single stranded, circular, or linear DNA. A biotinylated oligonucleotide comprising at least 15 nucleotides from the extended cDNA or 5' EST sequence is hybridized to the single stranded DNA. Hybrids between the biotinylated oligonucleotide and the single stranded DNA containing 30 the extended cDNA or EST sequence are isolated as described in Example 29 above. Thereafter, the single stranded DNA containing the extended cDNA or EST sequence is released from the beads and converted into double stranded DNA using a primer specific for the extended cDNA or 5' EST sequence or a primer corresponding to a sequence included in the cloning vector. The resulting double stranded DNA is transformed into bacteria. DNAs containing the 5' EST or extended cDNA sequences are identified by colony PCR or colony hybridization.

25

Once the upstream genomic sequences have been cloned and sequenced as described above, prospective promoters and transcription start sites within the upstream sequences may be identified by comparing the sequences upstream of the extended cDNAs or 5' ESTs with databases containing known transcription start sites, transcription factor binding sites, or promoter sequences.

In addition, promoters in the upstream sequences may be identified using promoter reporter vectors as described in Example 56.

EXAMPLE 56

Identification of Promoters in Cloned Upstream Sequences

The genomic sequences upstream of the extended cDNAs or 5' ESTs are cloned into a suitable promoter 10 reporter vector, such as the pSEAP-Basic, pSEAP-Enhancer, pβgal-Basic, pβgal-Enhancer, or pEGFP-1 Promoter Reporter vectors available from Clontech. Briefly, each of these promoter reporter vectors include multiple cloning sites positioned upstream of a reporter gene encoding a readily assayable protein such as secreted alkaline phosphatase, β galactosidase, or green fluorescent protein. The sequences upstream of the extended cDNAs or 5' ESTs are inserted into the cloning sites upstream of the reporter gene in both orientations and introduced into an appropriate host cell. The 15 level of reporter protein is assayed and compared to the level obtained from a vector which lacks an insert in the cloning site. The presence of an elevated expression level in the vector containing the insert with respect to the control vector indicates the presence of a promoter in the insert. If necessary, the upstream sequences can be cloned into vectors which contain an enhancer for augmenting transcription levels from weak promoter sequences. A significant level of expression above that observed with the vector lacking an insert indicates that a promoter sequence is present in the 20 inserted upstream sequence.

Appropriate host cells for the promoter reporter vectors may be chosen based on the results of the above described determination of expression patterns of the extended cDNAs and ESTs. For example, if the expression pattern analysis indicates that the mRNA corresponding to a particular extended cDNA or 5 $^{\prime}$ EST is expressed in fibroblasts, the promoter reporter vector may be introduced into a human fibroblast cell line.

Promoter sequences within the upstream genomic DNA may be further defined by constructing nested deletions in the upstream DNA using conventional techniques such as Exonuclease III digestion. The resulting deletion fragments can be inserted into the promoter reporter vector to determine whether the deletion has reduced or obliterated promoter activity. In this way, the boundaries of the promoters may be defined. If desired, potential individual regulatory sites within the promoter may be identified using site directed mutagenesis or linker scanning to obliterate 30 potential transcription factor binding sites within the promoter individually or in combination. The effects of these mutations on transcription levels may be determined by inserting the mutations into the cloning sites in the promoter reporter vectors.

EXAMPLE 57

Cloning and Identification of Promoters

Using the method described in Example 55 above with 5' ESTs, sequences upstream of several genes were obtained. Using the primer pairs GGG AAG ATG GAG ATA GTA TTG CCT G (SEQ ID NO:29) and CTG CCA TGT ACA TGA TAG AGA GAT TC (SEQ ID NO:30), the promoter having the internal designation P13H2 (SEQ ID NO:31) was obtained.

Using the primer pairs GTA CCA GGGG ACT GTG ACC ATT GC (SEQ ID NO:32) and CTG TGA CCA TTG CTC CCA AGA GAG (SEQ ID NO:33), the promoter having the internal designation P15B4 (SEQ ID NO:34) was obtained.

Using the primer pairs CTG GGA TGG AAG GCA CGG TA (SEQ ID NO:35) and GAG ACC ACA CAG CTA GAC AA (SEQ ID NO:36), the promoter having the internal designation P29B6 (SEQ ID NO:37) was obtained.

Figure 8 provides a schematic description of the promoters isolated and the way they are assembled with the corresponding 5' tags. The upstream sequences were screened for the presence of motifs resembling transcription factor binding sites or known transcription start sites using the computer program MatInspector release 2.0, August 1996.

Figure 9 describes the transcription factor binding sites present in each of these promoters. The columns labeled matrice provides the name of the MatInspector matrix used. The column labeled position provides the 5' postion of the promoter site. Numeration of the sequence starts from the transcription site as determined by matching the genomic sequence with the 5' EST sequence. The column labeled "orientation" indicates the DNA strand on which the site is found, with the + strand being the coding strand as determined by matching the genomic sequence with the sequence of the 5' EST. The column labeled "score" provides the MatInspector score found for this site. The column labeled "length" provides the length of the site in nucleotides. The column labeled "sequence" provides the sequence of the site found.

The promoters and other regulatory sequences located upstream of the extended cDNAs or 5' ESTs may be used to design expression vectors capable of directing the expression of an inserted gene in a desired spatial, temporal, developmental, or quantitative manner. A promoter capable of directing the desired spatial, temporal, developmental, and quantitative patterns may be selected using the results of the expression analysis described in Example 26 above. For example, if a promoter which confers a high level of expression in muscle is desired, the promoter sequence upstream of an extended cDNA or 5' EST derived from an mRNA which is expressed at a high level in muscle, as determined by the method of Example 26, may be used in the expression vector.

Preferably, the desired promoter is placed near multiple restriction sites to facilitate the cloning of the desired insert downstream of the promoter, such that the promoter is able to drive expression of the inserted gene. The promoter may be inserted in conventional nucleic acid backbones designed for extrachromosomal replication, integration into the host chromosomes or transient expression. Suitable backbones for the present expression vectors include retroviral backbones, backbones from eukaryotic episomes such as SV40 or Bovine Papilloma Virus, backbones from bacterial episomes, or artificial chromosomes.

30

Preferably, the expression vectors also include a polyA signal downstream of the multiple restriction sites for directing the polyadenylation of mRNA transcribed from the gene inserted into the expression vector.

Following the identification of promoter sequences using the procedures of Examples 55-57, proteins which interact with the promoter may be identified as described in Example 58 below.

EXAMPLE 58

Identification of Proteins Which Interact with Promoter Sequences, Upstream Regulatory Sequences, or mRNA

Sequences within the promoter region which are likely to bind transcription factors may be identified by homology to known transcription factor binding sites or through conventional mutagenesis or deletion analyses of reporter plasmids containing the promoter sequence. For example, deletions may be made in a reporter plasmid containing the promoter sequence of interest operably linked to an assayable reporter gene. The reporter plasmids carrying various deletions within the promoter region are transfected into an appropriate host cell and the effects of the deletions on expression levels is assessed. Transcription factor binding sites within the regions in which deletions reduce expression levels may be further localized using site directed mutagenesis, linker scanning analysis, or other techniques familiar to those skilled in the art. Nucleic acids encoding proteins which interact with sequences in the promoter may be identified using one-hybrid systems such as those described in the manual accompanying the Matchmaker One-Hybrid System kit avalilabe from Clontech (Catalog No. K1603-1). Briefly, the Matchmaker One-hybrid system is used as follows. The target sequence for which it is desired to identify binding proteins is cloned upstream of a selectable reporter gene and integrated into the yeast genome. Preferably, multiple copies of the target sequences are inserted into the reporter plasmid in tandem.

A library comprised of fusions between cDNAs to be evaluated for the ability to bind to the promoter and the activation domain of a yeast transcription factor, such as GAL4, is transformed into the yeast strain containing the integrated reporter sequence. The yeast are plated on selective media to select cells expressing the selectable marker linked to the promoter sequence. The colonies which grow on the selective media contain genes encoding proteins which bind the target sequence. The inserts in the genes encoding the fusion proteins are further characterized by sequencing. In addition, the inserts may be inserted into expression vectors or in vitro transcription vectors. Binding of the polypeptides encoded by the inserts to the promoter DNA may be confirmed by techniques familiar to those skilled in the art, such as gel shift analysis or DNAse protection analysis.

VII. Use of Extended cDNAs (or Genomic DNAs Obtainable Therefrom) in Gene Therapy

The present invention also comprises the use of extended cDNAs (or genomic DNAs obtainable therefrom) in gene therapy strategies, including antisense and triple helix strategies as described in Examples 57 and 58 below. In antisense approaches, nucleic acid sequences complementary to an mRNA are hybridized to the mRNA intracellularly, thereby blocking the expression of the protein encoded by the mRNA. The antisense sequences may prevent gene expression through a variety of mechanisms. For example, the antisense sequences may inhibit the ability of ribosomes

to translate the mRNA. Alternatively, the antisense sequences may block transport of the mRNA from the nucleus to the cytoplasm, thereby limiting the amount of mRNA available for translation. Another mechanism through which antisense sequences may inhibit gene expression is by interfering with mRNA splicing. In yet another strategy, the antisense nucleic acid may be incorporated in a ribozyme capable of specifically cleaving the target mRNA.

EXAMPLE 59

Preparation and Use of Antisense Oligonucleotides

The antisense nucleic acid molecules to be used in gene therapy may be either DNA or RNA sequences. They may comprise a sequence complementary to the sequence of the extended cDNA (or genomic DNA obtainable therefrom). The antisense nucleic acids should have a length and melting temperature sufficient to permit formation of an intracellular duplex having sufficient stability to inhibit the expression of the mRNA in the duplex. Strategies for designing antisense nucleic acids suitable for use in gene therapy are disclosed in Green et al., Ann. Rev. Biochem. 55:569-597 (1986) and Izant and Weintraub, Cell 36:1007-1015 (1984).

In some strategies, antisense molecules are obtained from a nucleotide sequence encoding a protein by reversing the orientation of the coding region with respect to a promoter so as to transcribe the opposite strand from that which is normally transcribed in the cell. The antisense molecules may be transcribed using in vitro transcription systems such as those which employ T7 or SP6 polymerase to generate the transcript. Another approach involves transcription of the antisense nucleic acids in vivo by operably linking DNA containing the antisense sequence to a promoter in an expression vector.

Alternatively, oligonucleotides which are complementary to the strand normally transcribed in the cell may be synthesized in vitro. Thus, the antisense nucleic acids are complementary to the corresponding mRNA and are capable of hybridizing to the mRNA to create a duplex. In some embodiments, the antisense sequences may contain modified sugar phosphate backbones to increase stability and make them less sensitive to RNase activity. Examples of modifications suitable for use in antisense strategies are described by Rossi et al., Pharmacol. Ther. 50(2):245-254, (1991).

Various types of antisense oligonucleotides complementary to the sequence of the extended cDNA (or genomic DNA obtainable therefrom) may be used. In one preferred embodiment, stable and semi-stable antisense oligonucleotides described in International Application No. PCT W094/23026 are used. In these molecules, the 3' end or both the 3' and 5' ends are engaged in intramolecular hydrogen bonding between complementary base pairs. These molecules are better able to withstand exonuclease attacks and exhibit increased stability compared to conventional antisense oligonucleotides.

In another preferred embodiment, the antisense oligodeoxynucleotides against herpes simplex virus types 1 and 2 described in International Application No. WO 95/04141.

In yet another preferred embodiment, the covalently cross-linked antisense oligonucleotides described in International Application No. WO 96/31523 are used. These double- or single-stranded oligonucleotides comprise one or more, respectively, inter- or intra-oligonucleotide covalent cross-linkages, wherein the linkage consists of an amide bond between a primary amine group of one strand and a carboxyl group of the other strand or of the same strand, respectively, the primary amine group being directly substituted in the 2' position of the strand nucleotide monosaccharide ring, and the carboxyl group being carried by an aliphatic spacer group substituted on a nucleotide or nucleotide analog of the other strand or the same strand, respectively.

The antisense oligodeoxynucleotides and oligonucleotides disclosed in International Application No. WO 92/18522 may also be used. These molecules are stable to degradation and contain at least one transcription control recognition sequence which binds to control proteins and are effective as decoys therefor. These molecules may contain "hairpin" structures, "dumbbell" structures, "modified dumbbell" structures, "cross-linked" decoy structures and "loop" structures.

In another preferred embodiment, the cyclic double-stranded oligonucleotides described in European Patent Application No. 0 572 287 A2 are used. These ligated oligonucleotide "dumbbells" contain the binding site for a transcription factor and inhibit expression of the gene under control of the transcription factor by sequestering the factor.

15

Use of the closed antisense oligonucleotides disclosed in International Application No. WO 92/19732 is also contemplated. Because these molecules have no free ends, they are more resistant to degradation by exonucleases than are conventional oligonucleotides. These oligonucleotides may be multifunctional, interacting with several regions which are not adjacent to the target mRNA.

The appropriate level of antisense nucleic acids required to inhibit gene expression may be determined using in vitro expression analysis. The antisense molecule may be introduced into the cells by diffusion, injection, infection or transfection using procedures known in the art. For example, the antisense nucleic acids can be introduced into the body as a bare or naked oligonucleotide, oligonucleotide encapsulated in lipid, oligonucleotide sequence encapsidated by viral protein, or as an oligonucleotide operably linked to a promoter contained in an expression vector. The expression vector may be any of a variety of expression vectors known in the art, including retroviral or viral vectors, vectors capable of extrachromosomal replication, or integrating vectors. The vectors may be DNA or RNA.

The antisense molecules are introduced onto cell samples at a number of different concentrations preferably between 1x10⁻¹⁰M to 1x10⁻⁴M. Once the minimum concentration that can adequately control gene expression is identified, the optimized dose is translated into a dosage suitable for use in vivo. For example, an inhibiting concentration in culture of 1x10⁻⁷ translates into a dose of approximately 0.6 mg/kg bodyweight. Levels of oligonucleotide approaching 100 mg/kg bodyweight or higher may be possible after testing the toxicity of the oligonucleotide in laboratory animals. It is additionally contemplated that cells from the vertebrate are removed, treated with the antisense oligonucleotide, and reintroduced into the vertebrate.

It is further contemplated that the antisense oligonucleotide sequence is incorporated into a ribozyme sequence to enable the antisense to specifically bind and cleave its target mRNA. For technical applications of ribozyme and antisense oligonucleotides see Rossi et al., *supra*.

In a preferred application of this invention, the polypeptide encoded by the gene is first identified, so that the

effectiveness of antisense inhibition on translation can be monitored using techniques that include but are not limited to
antibody-mediated tests such as RIAs and ELISA, functional assays, or radiolabeling.

The extended cDNAs of the present invention (or genomic DNAs obtainable therefrom) may also be used in gene therapy approaches based on intracellular triple helix formation. Triple helix oligonucleotides are used to inhibit transcription from a genome. They are particularly useful for studying alterations in cell activity as it is associated with a particular gene. The extended cDNAs (or genomic DNAs obtainable therefrom) of the present invention or, more preferably, a portion of those sequences, can be used to inhibit gene expression in individuals having diseases associated with expression of a particular gene. Similarly, a portion of the extended cDNA (or genomic DNA obtainable therefrom) can be used to study the effect of inhibiting transcription of a particular gene within a cell. Traditionally, homopurine sequences were considered the most useful for triple helix strategies. However, homopyrimidine sequences can also inhibit gene expression. Such homopyrimidine oligonucleotides bind to the major groove at homopurine:homopyrimidine sequences. Thus, both types of sequences from the extended cDNA or from the gene corresponding to the extended cDNA are contemplated within the scope of this invention.

EXAMPLE 60

Preparation and use of Triple Helix Probes

The sequences of the extended cDNAs (or genomic DNAs obtainable therefrom) are scanned to identify 10-mer to 20-mer homopyrimidine or homopurine stretches which could be used in triple-helix based strategies for inhibiting gene expression. Following identification of candidate homopyrimidine or homopurine stretches, their efficiency in inhibiting gene expression is assessed by introducing varying amounts of oligonucleotides containing the candidate sequences into tissue culture cells which normally express the target gene. The oligonucleotides may be prepared on an oligonucleotide synthesizer or they may be purchased commercially from a company specializing in custom oligonucleotide synthesis, such as GENSET, Paris, France.

The oligonucleotides may be introduced into the cells using a variety of methods known to those skilled in the art, including but not limited to calcium phosphate precipitation, DEAE-Dextran, electroporation, liposome-mediated transfection or native uptake.

Treated cells are monitored for altered cell function or reduced gene expression using techniques such as

Northern blotting, RNase protection assays, or PCR based strategies to monitor the transcription levels of the target
gene in cells which have been treated with the oligonucleotide. The cell functions to be monitored are predicted based
upon the homologies of the target gene corresponding to the extended cDNA from which the oligonucleotide was derived
with known gene sequences that have been associated with a particular function. The cell functions can also be

predicted based on the presence of abnormal physiologies within cells derived from individuals with a particular inherited disease, particularly when the extended cDNA is associated with the disease using techniques described in Example 53.

The oligonucleotides which are effective in inhibiting gene expression in tissue culture cells may then be introduced in vivo using the techniques described above and in Example 59 at a dosage calculated based on the in vitro results, as described in Example 59.

In some embodiments, the natural (beta) anomers of the oligonucleotide units can be replaced with alpha anomers to render the oligonucleotide more resistant to nucleases. Further, an intercalating agent such as ethidium bromide, or the like, can be attached to the 3' end of the alpha oligonucleotide to stabilize the triple helix. For information on the generation of oligonucleotides suitable for triple helix formation see Griffin et al. (Science 245:967-10 971 (1989).

EXAMPLE 61

Use of Extended cDNAs to Express an Encoded Protein in a Host Organism

The extended cDNAs of the present invention may also be used to express an encoded protein in a host organism to produce a beneficial effect. In such procedures, the encoded protein may be transiently expressed in the host organism or stably expressed in the host organism. The encoded protein may have any of the activities described above. The encoded protein may be a protein which the host organism lacks or, alternatively, the encoded protein may augment the existing levels of the protein in the host organism.

A full length extended cDNA encoding the signal peptide and the mature protein, or an extended cDNA encoding only the mature protein is introduced into the host organism. The extended cDNA may be introduced into the host organism using a variety of techniques known to those of skill in the art. For example, the extended cDNA may be injected into the host organism as naked DNA such that the encoded protein is expressed in the host organism, thereby producing a beneficial effect.

Alternatively, the extended cDNA may be cloned into an expression vector downstream of a promoter which is active in the host organism. The expression vector may be any of the expression vectors designed for use in gene therapy, including viral or retroviral vectors.

The expression vector may be directly introduced into the host organism such that the encoded protein is expressed in the host organism to produce a beneficial effect. In another approach, the expression vector may be introduced into cells in vitro. Cells containing the expression vector are thereafter selected and introduced into the host organism, where they express the encoded protein to produce a beneficial effect.

EXAMPLE 62

Use Of Signal Peptides Encoded By 5' Ests Or Sequences

Obtained Therefrom To Import Proteins Into Cells

The short core hydrophobic region (h) of signal peptides encoded by the 5'ESTS or extended cDNAs derived from the 5'ESTs of the present invention may also be used as a carrier to import a peptide or a protein of interest, so-

called cargo, into tissue culture cells (Lin et al., J. Biol. Chem., 270: 14225-14258 (1995); Du et al., J. Peptide Res., 51: 235-243 (1998); Rojas et al., Nature Biotech., 16: 370-375 (1998)).

When cell permeable peptides of limited size (approximately up to 25 amino acids) are to be translocated across cell membrane, chemical synthesis may be used in order to add the h region to either the C-terminus or the N-terminus to the cargo peptide of interest. Alternatively, when longer peptides or proteins are to be imported into cells, nucleic acids can be genetically engineered, using techniques familiar to those skilled in the art, in order to link the extended cDNA sequence encoding the h region to the 5' or the 3' end of a DNA sequence coding for a cargo polypeptide. Such genetically engineered nucleic acids are then translated either *in vitro* or *in vivo* after transfection into appropriate cells, using conventional techniques to produce the resulting cell permeable polypeptide. Suitable hosts cells are then simply incubated with the cell permeable polypeptide which is then translocated across the membrane.

This method may be applied to study diverse intracellular functions and cellular processes. For instance, it has been used to probe functionally relevant domains of intracellular proteins and to examine protein-protein interactions involved in signal transduction pathways (Lin et al., supra; Lin et al., J. Biol. Chem., 271: 5305-5308 (1996); Rojas et al., J. Biol. Chem., 271: 27456-27461 (1996); Liu et al., Proc. Natl. Acad. Sci. USA, 93: 11819-11824 (1996); Rojas et al., Bioch. Biophys. Res. Commun., 234: 675-680 (1997)).

Such techniques may be used in cellular therapy to import proteins producing therapeutic effects. For instance, cells isolated from a patient may be treated with imported therapeutic proteins and then re-introduced into the host organism.

Alternatively, the h region of signal peptides of the present invention could be used in combination with a nuclear localization signal to deliver nucleic acids into cell nucleus. Such oligonucleotides may be antisense oligonucleotides or oligonucleotides designed to form triple helixes, as described in examples 59 and 60 respectively, in order to inhibit processing and maturation of a target cellular RNA.

EXAMPLE 63

Reassembling & Resequencing of Clones

Full length cDNA clones obtained by the procedure described in Example 27 were double-sequenced. These sequences were assembled and the resulting consensus sequences were then reanalyzed. Open reading frames were reassigned following essentially the same process as the one described in Example 27.

After this reanalysis process a few abnormalities were revealed. The sequences presented in SEQ ID NOs: 47, 73, 79, 89, 91, 96, 126, 128, 134, and 139 are apparently unlikely to be genuine full length cDNAs. These clones are missing a stop codon and are thus more probably 3' truncated cDNA sequences. Similarly, the sequences presented in SEQ ID NOs: 45, 50, 54, 57, 73, 74, 89, 92, 95, 98, 126, 129, 130, 131 and 139 may also not be genuine full length cDNAs based on homology studies with existing protein sequences. Although both of these sequences encode a potential start methionine each could represent a 5' truncated cDNA.

In addition, SEQ ID NO: 115 was found to be an alternatively spliced transcript and the identities of some of the bases in SEQ ID NO: 131 were corrected.

Finally, after the reassignment of open reading frames for the clones, new open reading frames were chosen in some instances. For example, in the case of SEQ ID NOs: 41, 47, 50, 52, 54-56, 58, 59, 61, 74, 75, 79, 84, 89, 91, 92, 96, 98, 103, 105, 106, 126, 129, 131, and 133 the new open reading frames were no longer predicted to contain a signal peptide.

As discussed above, Table IV provides the sequence identification numbers of the extended cDNAs of the present invention, the locations of the full coding sequences in SEQ ID NOs: 40-140 and 242-377 (i.e. the nucleotides encoding both the signal peptide and the mature protein, listed under the heading FCS location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the signal peptides (listed under the heading SigPep Location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the mature proteins generated by cleavage of the signal peptides (listed under the heading Mature Polypeptide Location in Table IV), the locations in SEQ ID NOs: 40-140 and 242-377 of stop codons (listed under the heading Stop Codon Location in Table IV) the locations in SEQ ID NOs: 40-140 and 242-377 of polyA signals (listed under the heading g PolyA Signal Location in Table IV) and the locations of polyA sites (listed under the heading PolyA Site Location in Table IV).

As discussed above, Table V lists the sequence identification numbers of the polypeptides of SEQ ID NOs: 141-241 and 378-513, the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the full length polypeptide (second column), the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the signal peptides (third column), and the locations of the amino acid residues of SEQ ID NOs: 141-241 and 379-513 in the mature polypeptide created by cleaving the signal peptide from the fall length polypeptide (fourth column). In Table V, and in the appended sequence listing, the first amino acid of the mature protein resulting from cleavage of the signal peptide is designated as amino acid number 1 and the first amino acid of the signal peptide is designated with the appropriate negative number, in accordance with the regulations governing sequence listings.

25

EXAMPLE 64

Functional Analysis of Predicted Protein Sequences

Following double-sequencing, new contigs were assembled for each of the extended cDNAs of the present invention and each was compared to known sequences available at the time of filing. These sequences originate from the following databases: Genbank (release 108 and daily releases up to October, 15, 1998), Genseq (release 32) PIR (release 33) and SwissProt (release 35). The predicted proteins of the present invention matching known proteins were further classified into 3 categories depending on the level of homology.

The first category contains proteins of the present invention exhibiting more than 70% identical amino acid residues on the whole length of the matched protein. They are clearly close homologues which most probably have the same function or a very similar function as the matched protein.

The second category contains proteins of the present invention exhibiting more remote homologies (40 to 70% over the whole protein) indicating that the protein of the present inventionmay have functions similar to those of the homologous protein.

The third category contains proteins exhibiting homology (90 to 100%) to a domain of a known protein indicating that the matched protein and the protein of the invention may share similar features.

It should be noted that the numbering of amino acids in the protein sequences discussed in Figures 10 to 15, and Table VIII, the first methionine encountered is designated as amino acid number 1. In the appended sequence listing, the first amino acid of the mature protein resulting from cleavage of the signal peptide is designated as amino acid number 1, and the first amino acid of the signal peptide is designated with the appropriate negative number, in accordance with the regulations governing sequence listings.

In addition all of the corrected amino acid sequences (SEQ ED NOs: 141-241 and 378-513) were scanned for the presence of known protein signatures and motifs. This search was performed against the Prosite 15.0 database, using the Proscan software from the GCG package- Functional signatures and their locations are indicated in Table VIII.

15 A) Proteins which are closely related to known proteins

Protein of SEQ ID NO: 217

The protein of SEQ ID NO: 217 encoded by the extended cDNA SEQ ID NO: 116 isolated from lymphocyte shows complete identity to a human protein TFAR19 that may play a role in apoptosis (Genbank accession number AF014955, SEQ ID NO: 516) as shown by the alignment in figure 10.

Taken together, these data suggest that the protein of SEQ ID NO: 217 may be involved in the control of development and homeostasis. Thus, this protein may be useful in diagnosis and/or treating several types of disorders including, but not limited to, cancer, autoimmune disorders, viral infections such as AIDS, neurodegenerative disorders, osteoporosis.

25 Proteins of SEQ ID NOs: 174, 175 and 232

The proteins of SEQ ID NOs: 174, 175 and 232 encoded by the extended cDNAs SEQ ID NOs:. 73, 74 and 131 respectively and isolated from lymphocytes shows complete extensive homologies to a human secreted protein (Genseq accession number W36955, SEQ ID NO: 517). As shown by the alignments of figure 11, the amino acid residues are identical to those of the 110 amino acid long matched protein except for positions 51 and 108-110 of the matched protein for the protein of SEQ ID NOs: 174, for positions 48, 94 and 108-110 of the matched protein of SEQ ID NOs:175 and for positions 94, and 108-110 of the matched protein for the protein of SEQ ID NOs: 232. Proteins of SEQ ID NOs: 174 and 232 may represent alternative forms issued from alternative use of polyadenylation signals.

Taken together, these data suggest that the proteins of SEO ID NOs: 174, 175 and 232 may play a role in cell proliferation and/or differentiation, in immune responses and/or in haematopoeisis. Thus, this protein or part therein,

may be useful in diagnosing and treating several disorders including, but not limited to, cancer, immunological, haematological and/or inflammatory disorders. It may also be useful in modulating the immune and inflammatory responses to infectious agents and/or to suppress graft rejection.

5 Proteins of SEQ ID NO: 231

The protein of SEQ ID NO: 231 encoded by the extended cDNA SEQ ID NO: 130 shows extensive homology with the human E25 protein (Genbank accession number AF038953, SEQ ID NO: 515). As shown by the alignments in figure 12, the amino acid residues are identical except for position 159 in the 263 amino acid long matched sequence. The matched protein might be involved in the development and differentiation of haematopoietic stem/progenitor cells.

10 In addition, it is the human homologue of a murine protein thought to be involved in chondro-osteogenic differentiation and belonging to a novel multigene family of integral membrane proteins (Deleersnijder et al, J. Biol. Chem., 271: 19475-19482 (1996)).

The protein of invention contains two short segments from positions 1 to 21 and from 100 to 120 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, 10: 685-686 (1994)). The first transmembrane domains matches exactly those predicted for the murine E25 protein.

Taken together, these data suggest that the protein of SEQ ID NO: 231 may be involved in cellular proliferation and differentiation. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer and embryogenesis disorders.

20 Protein of SEQ ID NO: 196

The protein of SEQ ID NO: 196 encoded by the extended cDNA SEQ ID NO: 95 shows extensive homology with the human seventransmembrane protein (Genbank accession number Y11395, SEQ ID NO: 518) and its murine homologue (Genbank accession number Y11550). As shown by the alignments in figure 13, the amino acid residues are identical except for position 174 in the 399 amino acid long human matched sequence. The matched protein potentially associated to stomatin may act as a G-protein coupled receptor and is likely to be important for the signal transduction in neurons and haematopoietic cells (Mayer et al, Biochem. Biophys. Acta., 1395: 301-308 (1998)).

Taken together, these data suggest that the protein of SEQ ID NOs: 196 may be involved in signal transduction. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases cardiovascular disorders, hypertension, renal injury and repair and septic shock.

Protein of SEQ ID NO: 158

The protein of SEQ ID NOs: 158 encoded by the extended cDNA SEQ ID NO: 57 shows homology with the murine subunit 7a of the COP9 complex (Genbank accession number AF071316, SEQ ID NO: 520). As shown by the

alignments in figure 14, the amino acid residues are identical except for positions 90, 172 and 247 in the 275 amino acid long matched sequence. This complex is highly conserved between mammals and higher plants where it has been shown to act as a repressor of photomorphogenesis All the components of the mammalian COP9 complex contain structural features also present in components of the proteasome regulatory complex and the translation initiation complex eIF3 complex, suggesting that the mammalian COP9 complex is an important cellular regulator modulating multiple signaling pathways (Wei et al, Curr. Biol., 8: 919-922 (1998)).

Taken together, these data suggest that the protein of SEQ ID NO: 158 may be involved in cellular signaling, probably as a subunit of the human COP9 complex. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, cardiovascular disorders, hypertension, renal injury and repair and septic shock.

Protein of SEQ ID NO: 226

The protein of SEQ ID NO: 226 encoded by the extended cDNA SEQ ID NO: 125 shows homology with the bovine subunit B14.5B of the NADH-ubiquinone oxidureductase complex (Arizmendi et al, FEBS Lett., 313: 80-84 (1992) and Swissprot accession -number Q02827, SEQ ID NO: 514). As shown by the alignments in figure 15, the amino acid residues are identical except for positions 3-4, 6-12, 32-34, 47, 53-55, 67 and 69-74 in the 120 amino acid long matched sequence. This complex is the first of four complexes located in the inner mitochondrial membrane and composing the mitochondrial electron transport chain. Complex I is involved in the dehydrogenation of NADH and the transportation of electrons to coenzyme Q. It is composed of 7 subunits encoded by the mitochondrial genome and 34 subunits encoded by the nuclear genome. It is also thought to play a role in the regulation of apoptosis and necrosis. Mitochondriocytopathies due to complex I deficiency are frequently encountered and affect tissues with a high energy demand such as brain (mental retardation, convulsions, movement disorders), heart (cardiomyopathy, conduction disorders), kidney (Fanconi syndrome), skeletal muscle (exercise intolerance, muscle weakness, hypotonia) and/or eye (opthmaloplegia, ptosis, cataract and retinopathy). For a review on complex I see Smeitink et al., Hum. Mol. Gent., 7: 1573-1579 (1998).

Taken together, these data suggest that the protein of SEQ ID NO: 226 may be part of the mitochondrial energy-generating system, probably as a subunit of the NADH-ubiquinone oxidoreductase complex. Thus, this protein or part therein, may be useful in diagnosing and/or treating several disorders including, but not limited to, brain disorders (mental retardation, convulsions, movement disorders), 'heart disorders (cardiomyopathy, conduction disorders), kidney disorders (Fanconi syndrome), skeletal muscle disorders (exercise intolerance, muscle weakness, hypotonia) and/or eye disorders opthmalmoplegia, ptosis, cataract and retinopathy).

B) Proteins which are remotely related to proteins with known functions

<u>Proteins of SEQ ID NOs: 149, 150 and 211</u>

The proteins of SEQ ID NOs: 1.49,150 and 211 encoded by the extended cDNAs SEQ ID NOs: 48, 49 and 110 respectively and found in, skeletal muscle shows homologies with T1/ST2 ligand polypeptide of either human (Genbank accession number U41804 and Genseq accession number W09639) or rodent species (Genbank accession number U41805 and Genseq accession number W09640). These polypeptides are thought to be cytokines that bind to the ST2 receptor, a member of the immunoglobulin family homologous to the interleukin-1 receptor and present on some lymphoma cells. They are predicted to be cell-surface proteins containing a short transmembrane domain. (Gayle *et al, J. Biol. Chem.*, 271: 5784-5789 (1996)). Proteins of SEQ ID NOs: 149, 150 and 211 may represent alternative forms issued from alternative use of polyadenylation signals.

The protein of invention contains two short transmembrane segments from positions 5 to 25 and from 195 to 215 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, **10**:685-686 (1994)). The second transmembrane domain matches exactly those of the matched cell-surface protein.

Taken together, these data suggest that the protein of SEQ ID NOs: 149, 150 and 211 may act as a cytokine, thus may play a role in the regulation of cell growth and differentiation and/or in the regulation of the immune response. Thus, this protein or part therein, may be useful in diagnosing and treating several disorders including, but not limited to, cancer, immunological, haematological and/or inflammatory disorders. It may also be useful in modulating the immune and inflammatory responses to infectious agents such as HIV and/or to suppress graft rejection.

Protein of SEQ ID NO: 177

The protein SEQ ID NO: 177 found in testis encoded by the extended cDNA SEQ ID NO: 76 shows homologies to serine protease inhibitor proteins belonging to the pancreatic trypsin inhibitor family (Kunitz) such as the extracellular proteinase inhibitor named chelonianin (Swissprot accession number P00993). The characteristic PROSITE signature of this family is conserved in the protein of the invention (positions 69 to 87) except for a drastic change of the last cysteine residue into an arginine residue.

Taken together, these data suggest that the protein of SEQ ID NO: 177 may be a protease inhibitor, probably of the Kunitz family. Thus, this protein or part therein, may be useful in diagnosing and treating several disorders including but not limited to, cancer and neurodegenerative disorders such as Alzheimer's disease.

Protein of SEQ ID NO: 146

The protein SEQ ID NO: 146 encoded by the extended cDNA SEQ ID NO: 45 shows homology to human apolipoprotein L (Genbank accession number AF019225). The matched protein is a secreted high density lipoprotein associated with apoA-I-containing lipoproteins which play a key role in reverse cholesterol transport.

Taken together, these data suggest that the protein of SEQ ID NO. 146 may play a role in lipid metabolism. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to,

hyperlipidemia, hypercholesterolemia, atherosclerosis, cardiovascular disorders such as, coronary heart disease, and neurodegenerative disorders such as Alzheimer's disease or dementia.

Protein of SEQ ID NO: 163

5

The protein SEQ ED NO: 163 encoded by the extended cDNA SEQ ID NO: 62 shows homology to the yeast autophagocytosis protein AUT1 (SwissProt accession number P40344). The matched protein is required for starvation-induced non-specific bulk transport of cytoplasmic proteins to the vacuole.

Taken together, these data suggest that the protein of SEQ ID NO: 163 may play a role in protein transport.

Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to,
autoimmune disorders and immune disorders due to dysfunction of antigen presentation.

C) Proteins homologous to a domain of a protein with known function

Protein of SEQ ID NO: 214

The protein of SEQ ID NO: 214 encoded by the extended cDNA SEQ ID NO: 113 and expressed in adult brain shows extensive homology to part of the murine SHYC protein (Genbank accession number AF072697) which is expressed in the developing and embryonic nervous system as well as along the olfactory pathway in adult brains (Köster et al., Neuroscience Letters., 252: 69-71 (1998)).

Taken together, these data suggest that the protein of SEQ ID NO: 214 may play a role in nervous system development and function. Thus, this protein may be useful in diagnosing and/or treating cancer and/or brain disorders, including neurodegenerative disorders such as Alzheimer's and Parkinson's diseases.

Protein of SEQ ID NO: 225

The protein of SEQ ID NO: 225 encoded by the extended cDNA SEQ ID NO: 124 and expressed in adult prostate belong to the phosphatidylethanolainin-binding protein from which it exhibits the characteristic PROSITE signature from positions 90 to 112 (see table VIII). Proteins from this widespread family, from nematodes to fly, yeast, rodent and primate species, bind hydrophobic ligands such as phospholipids and nucleotides. They are mostly expressed in brain and in testis and are thought to play a role in cell growth and/or maturation, in regulation of the sperm maturation, motility and 'in membrane remodeling. They may act either through signal transduction or through oxidoreduction reactions (for a review see Schoentgen and Jollès, *FEBS Letters*, 369 : 22-26 (1995)).

Taken together, these data suggest that the protein of SEQ ID NO: 225 may play a role in cell. Thus, these growth, maturation and in membrane remodeling and/or may be related to male fertility. Thus, this protein may be useful in diagnosing and/or treating cancer, neurodegenerative diseases, and/of, disorders related to male fertility and sterility.

Protein of SEQ ID NO: 153

30

The protein of SEQ ID NO: 153 encoded by the extended cDNA SEQ ID NO. 52 and expressed in brain exhibits homology to different integral membrane proteins. These membrane proteins include the nematode protein SRE-2 (Swissprot accession number Q09273) that belongs to the multigene SRE family of *C. elegans* receptor-like proteins and a family of tricarboxylate carriers conserved between flies and mammals. One member of this matched family is the rat tricarboxylate carrier (Genbank accession number S70011), an anion transporter localized in the inner membrane of mitochondria and involved in the biosynthesis of fatty acids and cholesterol. The protein of the invention contains a short transmembrane segments from positions 5 to 25 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, 10:685-686 (1994)).

Taken together, these data suggest that the protein of SEO ID NO: 153 may play a role in signal transduction and/or in molecule transport. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, immune disorders, cardiovascular disorders, hypertension, renal injury and repair and septic shock

Protein of SEQ ID NO: 213

15

20

25

The protein of SEQ ID NO: 213 encoded by the extended cDNA SEQ ID NO: 112 and expressed in brain exhibits homology with part of the tRNA pseudouridine 55 synthase found in *Escherichia Coli* (Swissprot accession number P09171). This bacterial protein belongs to the NAP57/CBF5/TRUB family of nucleolar proteins found in bacteria, yeasts and mammals involved in rRNA or tRNA biosynthesis, ribosomal subunit assembly and/or centromere/mircotubule binding.

Taken together, these data suggest that the protein of SEQ ID NO: 213 may play a role in rRNA or tRNA biogensis and function. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, hearing loss or disorders linked to chromosomal instability such as dyskeratosis.

Protein of SEQ ED NO: 240

The protein of SEQ ID NO: 240 encoded by the extended cDNA SEQ ID NO: 139 and expressed in brain exhibits homology with a family of eukaryotic cell surface antigens containing 4 transmembrane domains. The PROSITE signature for this family is conserved in the protein of the invention except for a substitution of an alanine residue in place of any of the following hydrophic residues: leucine, valine, isoleucine or methionine (positions 21 to 36).

The protein of the invention contains three short transmembrane segments from positions 6 to 26, 32 to 52 and from 56 to 76 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, 10: 685-686 (1994)). These transmembrane domains match the last three transmembrane domains of the matched protein family.

Taken together, these data suggest that the protein of SEQ ID NO: 240 may play a role in immunological and/or inflammatory responses, probably as a cell surface antigen. Thus, this protein or part therein, may be useful in diagnosing and treating several disorders including, but not limited to, cancer, immunological, haematological and/or

inflammatory disorders. It may also be useful in modulating the immune and inflammatory responses to infectious agents and/or to suppress graft rejection.

Protein of SEQ ID NO: 239

5

10

20

The protein of SEQ ID NO: 239 encoded by the extended cDNA SEQ ID NO: 138 exhibits homology with a conserved region in a family of NA+/H+ exchanger conserved in yeast, nematode and mammals. These cation/proton exchangers are integral membrane proteins with 5 transmembrane segments involved in intracellular pH regulation, maintenance of cell volume, reabsorption of sodium across specialized epithelia, vectorial transport and are also thought to play a role in signal transduction and especially in the induction of cell proliferation and in the induction of apoptosis.

The protein of invention contains four short transmembrane segments from positions 21 to 41, 48 to 68 and from 131 to 151 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, 10: 685-686 (1994)). The third and fourth transmembrane domains match the fourth and fifth transmembrane segments of the matched family of proteins.

Taken together, these data suggest that the protein of SEQ ID NO: 239 may play a role in membrane

15 permeability and/or in signal transduction. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, cardiovascular disorders, hypertension, renal injury and repair, septic shock as well as disorders of membrane permeability such as diarrhea.

Protein of SEQ ID NO: 200

The protein of SEQ ID NO: 200 encoded by the extended cDNA SEQ ED NO: 99 and expressed in brain exhibits extensive homology to the N-terminus of cell division cycle protein 23 (Genbank accession number AF053977) and also to a lesser extent to its homologue in *Saccharomyces cerevisiae*. The matched protein is required for chromosome segregation and is part of the anaphae-promoting complex necessary for cell cycle progression to mitosis.

Taken together, these data suggest that the protein of SEQ ID NO: 200 may play a role in cellular mitosis.

Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer and leukemia.

Protein of SEQ ID NO: 230

The protein of SEQ ID NO: 230 encoded by the extended cDNA SEQ ID NO: 129 exhibits extensive homology to the C-terminus of the eta subunit of T-complex polypeptide 1 conserved from yeasts to mammals, and even complete identity with the last 54 amino acid residues of the human protein (Genbank accession number AF026292). The matched protein is a chaperonin which assists the folding of actins and tubulins in eukaryotic cells upon ATP hydrolysis.

Taken together, these data suggest that the protein of SEQ ID NO: 230 may play a role in the folding, transport, assembly and degradation of proteins. Thus, this protein may be useful in diagnosing and/or treating several

types of disorders including, but not limited to, cancer, cardiovascular disorders, immune disorders, neurodegenerative disorders, osteoporosis and arthritis.

Protein of SEQ ED NO: 167

5

The protein of SEQ ID NO: 167 encoded by the extended cDNA SEQ ID NO: 66 exhibits homology to a monkey pepsinogen A-4 precursor (Swissprot accession number P27678) and to related members of the aspartyl protease family. The matched protein belongs to a family of widely distributed proteolytic enzymes known to exist in vertebrate, fungi, plants, retroviruses and some plant viruses.

Taken together, these data suggest that the protein of SEQ ID NO: 167 may play a role in the degradation of proteins. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, autoimmune disorders and immune disorders due to dysfunction of antigen presentation.

Protein of SEQ ID NO: 179

The protein of SEQ ID NO: 179 encoded by the extended cDNA SEQ ID NO: 78 found in testis exhibits

15 homology to part of mammalian collipase precursors. Collipases are secreted cofactors for pancreatic lipases that allow the lipase to anchor at the water-lipid interface. Collipase plays a crucial role in the intestinal digestion and absorption of dietary fats. The 5 cysteines characteristic for this protein family are conserved in the protein of the invention although the collipase PROSITE signature is not.

Taken together, these data suggest that the protein of SEQ ED NO: 179 may play a role in the lipid metabolism and/or in male fertility. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, hyperlipidemia, hypercholesterolemia, atherosclerosis, cardiovascular disorders such as coronary heart disease, and neurodegenerative disorders such as Alzheimer's disease or dementia, and disorders linked to male fertility.

25 Protein of SEQ ID NO: 227

The protein of SEQ ID NO: 227 encoded by the extended cDNA SEQ ID NO: 126 exhibits extensive homology to the ATP binding region of a whole family of serine/threonine protein kinases belonging to the CDC2/CDC28 subfamily.

The PROSITE signature characteristic for this domain is present in the protein of the invention from positions 10 to 34.

Taken together, these data suggest that the protein of SEQ ED NO: 158 may bind ATP, and even be a protein 30 kinase. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, cardiovascular disorders, hypertension, renal injury and repair and septic shock.

Although this invention has been described in terms of certain preferred embodiments, other embodiments which will be apparent to those of ordinary skill in the art in view of the disclosure herein are also within the scope of this invention. Accordingly, the scope of the invention is intended to be defined only by reference to the appended claims.

As discussed above, the extended cDNAs of the present invention or portions thereof can be used for various purposes. The polynucleotides can be used to express recombinant protein for analysis, characterization or therapeutic use; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in disease states); as molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to identify chromosomes or to map related gene positions; to 10 compare with endogenous DNA sequences in patients to identify potential genetic disorders; as probes to hybridize and thus discover novel, related DNA sequences; as a source of information to derive PCR primers for genetic fingerprinting; for selecting and making oligomers for attachment to a "gene chip" or other support, including for examination for expression patterns; to raise anti-protein antibodies using DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein which binds or 15 potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the polynucleotide can also be used in interaction trap assays (such as, for example, that described in Gyuris et al., Cell 75:791-803 (1993)) to identify polynucleotides encoding the other protein with which binding occurs or to identify inhibitors of the binding interaction.

The proteins or polypeptides provided by the present invention can similarly be used in assays to determine biological activity, including in a panel of multiple proteins for high-throughput screening; to raise antibodies or to elicit 20 another immune response; as a reagent (including the labeled reagent) in assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other 25 protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of being developed into reagent grade or kit format for commercialization as research products.

Methods for performing the uses listed above are well known to those skilled in the art. References disclosing 30 such methods include without limitation "Molecular Cloning; A Laboratory Manual", 2d ed., Cole Spring Harbor Laboratory Press, Sambrook, J., E.F. Fritsch and T. Maniatis eds., 1989, and "Methods in Enzymology; Guide to Molecular Cloning Techniques", Academic Press, Berger, S.L. and A.R. Kimmel eds., 1987.

Polynucleotides and proteins of the present invention can also be used as nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid supplement, use as a carbon source, use as a

WO 99/31236 PCT/IB98/02122

-98-

nitrogen source and use as a source of carbohydrate. In such cases the protein or polynucleotide of the invention can be added to the feed of a particular organism or can be administered as a separate solid or liquid preparation, such as in the form of powder, pills, solutions, suspensions or capsules. In the case of microorganisms, the protein or polynucleotide of the invention can be added to the medium in or on which the microorganism is cultured.

SEQUENCE LISTING FREE TEXT

The following free text appears in the accompanying Sequence Listing: In vitro transcription product oligonucleotide

5 promoter
transcription start site
Von Heijne matrix
Score

matinspector prediction

10 name

TABLE I

SEQ ID NO. in Present application	Provisional Application Disclosing Sequence	SEQ ID NO. in provisional application
40	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	51
41	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	72
42	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	52
43	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	78
44	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	73
45	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	41
46	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	67
47	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	82
48	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	80
49	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	81
50	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	53
· 51	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	54
52	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	195
53	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	44
54	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	46
55	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	68
56	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	48
57	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	55
58	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	49
59	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	50
60	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	97
61	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	51
62	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	69
63	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	49
64	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	199
65	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	53
66	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	57
67	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	54
68	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	55
69	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	58
70	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	59

CONT. TABLE

CONT. TABLE I		٠.
71	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	60
72	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	112
73	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	52
74	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	59
75	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	60
76	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	136
77	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	75
78	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	61
79	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	61
80	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	130
81	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	65
82	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	54
83	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	78
84	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	63
85	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	65
86	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	152
87	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	66
88	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	67
89	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	60
90	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	68
91	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	61
92	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	62
93	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	166
94	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	70
95	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	73
96	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	63
97	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	52
98	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	62
99	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	176
100	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998,	63
101	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	187
102	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	190
103	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	83
104	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	180
105	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	64
106	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	69

CONT TARIF

CONT. TABLE I		٠.
107	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	40
108	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	77
109	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	43
110	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	82
111	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	76
112	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	43
113	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	46
114	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	47
115	U.S. Provisional Patent Application Serial No. 60/066,677, filed Nov. 13, 1997	53
116	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	58
117	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	74
118	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	71
119	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	145
120	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	67
121	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	58
122	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	72
123	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	73
124	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	70
125	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	40
126	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	44
127	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	45
128	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	47
129	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	48
130	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	51
131	U.S. Provisional Patent Application Serial No. 60/066,677, filed Nov. 13, 1997	50
132	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	56
133	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	57
134	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	71
135	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	72
136	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	64
137	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	65
138	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	66
139	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	74
140	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	67
242	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	75
243	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	76

CONT. TABLE I		·.
244	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	77
245	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	78
246	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	79
247	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	80
248	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	81
249	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	82
250	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	83
251	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	84
252	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	85
253	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	86
254	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	87
255	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	88
256	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	89
257	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	90
258	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	91
259	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	92
260	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	93
261	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	94
262	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	95
263	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	96
264	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	97
265	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	98
266	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	99
267	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	100
268	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	101
269	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	102
270	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	103
271	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	104
272	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	105
273	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	106
274	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	107
275	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	108
276	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	109
277	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	110
278	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	111
279	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	112

CONT. TABLE

CONT. TABLE I		
280	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	113
281	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	114
282	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	115
283	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	116
284	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	117
285	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	118
286	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	119
287	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	120
288	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	121
289	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	122
290	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	123
291	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	124
292	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	125
293	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	126
294	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	127
295	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	128
296	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	129
297	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	130
298	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	131
299	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	132
300	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	133
301	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	134
302	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	135
303	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	136
304	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	137
305	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	138
306	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	139
307	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	140
308	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	141
309	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	142
310	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	143
311	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	. 144
312	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	145
313	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	146
314	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	147
315	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	148

CONT TARIFI

CONT. TABLE I		·.
316	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	149
317	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	150
318	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	151
319	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	152
320	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	153
321	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	154
322	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	155
323	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	156
324	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	157
325	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	158
326	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	159
327	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	160
328	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	161
329	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	162
330	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	163
331	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	164
332	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	165
333	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	166
334	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	167
335	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	168
336	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	169
337	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	170
338	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	171
339	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	172
340	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	173
341	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	174
342	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	175
343	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	176
344	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	177
345	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	178
346	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	179
347	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	180
348	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	181
349	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	182
350	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	183
351	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	184

-106-

CONT. TABLE I

	••
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	185
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	186
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	187
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	188
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	189
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	190
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	191
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	192
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	193
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	194
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	195
L.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	196
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	197
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	1998
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	199
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	200
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	201
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	202
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	203
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	204
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	205
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	206
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	207
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	208
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	209
U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	210
	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997 U.S. Provisional Patent Application Serial No.

TABLE II: Parameters used for each step of EST analysis

		Search Charac	teristics	Selection Charac	teristics
Step	Program	Strand	Parameters	Identity (%))	Length (bp)
Miscellaneous	Blastn	both	S-61 X-16	90	17
tRNA	Fasta	both	•	80	60
rRNA	Blastn	both	S=108	80	40
mtRNA	Blastn	both	S=108	80	40
Procaryotic	Blastn	both	S-144	90	40
Fungal	Blastn	both	S=144	90	40
Alu	fasta*	both	•	70	40
L1	Blastn	both	S=72	70	40
Repeats	Blastn	both	S=72	70	40
Promoters	Blastn	top	S-54 X-16	90	15⊥
Vertebrate	fasta*	both	S-108	90	30
ESTs	Blatsn	both	S-108 X-16	90	30
Proteins	blastxn	top	E-0.001		

^{*} use "Quick Fast" Database Scanner

 $[\]pm$ alignment further constrained to begin closer than 10bp to EST\5' end 5 $~\eta~$ using BLOSUM62 substitution matrix

TABLE III: Parameters used for each step of extended cDNA analysis

	Search characte	ristics		Selection	n characteristic	CS	
Step	Program	Strand	Parameters	Identity (%)	Length (bp)	Comments	
miscellaneous •	FASTA	both	•	90	15		
tRNA*	FASTA	both	•	80	90		
rRNA ¹	BLASTN	both	S-108	80	40		
mtRNA*	BLASTN	both	S-108	80	40		
Procaryotic ¹	BLASTN	both	S-144	90	40		
Fungal*	BLASTN	both	S-144	90	40		
Alu*	BLASTN	both	S-72	70	40	max 5 matches, masking	
L1 ¹	BLASTN	both	S-72	70	40	max 5 matches, masking	
Repeats*	BLASTN	both	S=72	70	40	masking	
PolyA	BLAST2N	top	W-6,S-10,E-1000	90	8	in the last 20 nucleotides	
Polyadenylati on signal	•	top	AATAAA allowing 1 mis	smatch		in the 50 nucleotides preceding the 5' end of the polA	
Vertebrate*	BLASTN then FASTA	both		90 then 70	30	first BLASTN and then FASTA on matching sequences	
ESTs*	BLAST2N	both	•	90	30		
Geneseq	BLASTN	both	W-8, B-10	90	30		
ORF	BLASTP	top	W-8, B-10		•	on ORF proteins, max 10 matches	
Proteins*	BLASTX	top	E=0.001	70	30		

 $^{^{\}rm t}$ steps common to EST analysis and using the same algorithms and parameters $^{\rm t}$ steps also used in EST analysis but with different algorithms and/or parameters

TABLE IV

			IA	BLE IV		
ld	FCS Location	SigPep Location	Mature Polypeptide Location	Stop Codon Location	PolyA Signal Location	PolyA Site Location
40	7 through 471	7 through 99	100 through 471	472	537 through 542	554 through 568
41	168 through 332	-	168 through 332	333	557 through 562	•
42	51 through 251	51 through 110	111 through 251	252	849 through 854	882 through 895
43	20 through 613	20 through 82	83 through 613	614		
44	12 through 416	12 through 86	87 through 416	417	425 through 430	445 through 458
45	276 through 1040	276 through 485	486 through 1040	1041		2024 through 2036
46	443 through 619	443 through 589	590 through 619	620		1267 through 1276
47	206 through 747	•	206 through 747	1.	1.	·
48	36 through 521	36 through 104	105 through 521	522	528 through 533	548 through 561
49	36 through 395	36 through 104	105 through 395	396	599 through 604	619 through 632
50	21 through 41		21 through 41	42	328 through 333	357 through 370
51	35 through 631	35 through 160	161 through 631	632	901 through 906	979 through 994
52	271 through 399		271 through 399	400		
53	103 through 252	103 through 213	214 through 252	253		588 through 597
54	2 through 460	·	2 through 460	461	713 through 718	735 through 748
55	31 through 231	·	31 through 231	232	769 through 774	690 through 703
56	305 through 565		305 through 565	566	694 through 699	713 through 725
57	124 through 873	124 through 378	379 through 873	874	1673 through 1678	1694 through 1705
58	135 through 206	•	135 through 206	207	850 through 855	1056 through 1069
59	135 through 818	·	135 through 818	819	909 through 914	1071 through 1084
60	33 through 290	33 through 92	93 through 290	291		
61	485 through 616		485 through 616	617		669 through 682
62	54 through 995	54 through 227	228 through 995	996	1130 through 1135	1181 through 1191
63	657 through 923	657 through 896	897 through 923	924	957 through 962	974 through 1008
64	18 through 311	18 through 62	63 through 311	312		
65	151 through 426	151 through 258	259 through 426	427	505 through 510	527 through 538
66	10 through 1062	10 through 57	58 through 1062	1063	1710 through 1715	1735 through 1747
67	78 through 491	78 through 218	219 through 491	492	1652 through 1657	1673 through 1686
68	69 through 371	69 through 287	288 through 371	372	510 through 515	530 through 542
69	2 through 757	2 through 205	206 through 757	758	•	1160 through 1174
70	2 through 1051	2 through 205	206 through 1051	1052	1248 through 1253	1272 through 1285
71	2 through 1171	2 through 205	206 through 1171	1172	1368 through 1373	1386 through 1398
72	42 through 611	42 through 287	288 through 611	612	787 through 792	808 through 821
73	62 through 916	62 through 757	758 through 916			904 through 916
74	62 through 520	•	62 through 520	521	1124 through 1129	1141 through 1153
75	21 through 167	•	21 through 167	168	•	
76	22 through 318	22 through 93	94 through 318	319	497 through 502	516 through 526
77	8 through 292	8 through 118	119 through 292	293	317 through 322	339 through 352
78	16 through 378	16 through 84	85 through 378	379	502 through 507	522 through 542

CONT. TABLE IV

79						
	57 through 233	·	57 through 233		•	•
80	83 through 340	83 through 124	125 through 340	341	573 through 578	607 through 660
81	47 through 541	47 through 220	221 through 541	542	•	597 through 605
82	46 through 285	46 through 150	151 through 285	286	364 through 369	385 through 396
83	22 through 240	22 through 84	85 through 240	241	397 through 402	421 through 432
84	89 through 382	•	89 through 382	383	1	408 through 420
85	80 through 415	80 through 142	143 through 415	416	471 through 476	488 through 501
86	152 through 361	152 through 283	284 through 361	362	·	•
87	32 through 307	32 through 70	71 through 307	308	1240 through 1245	1261 through 1272
88	114 through 734	114 through 239	240 through 734	735	768 through 773	793 through 804
89	199 through 802	•	199 through 802		780 through 785	791 through 802
90	38 through 1174	38 through 148	149 through 1174	1175	1452 through 1457	1478 through 1490
91	26 through 361	·	26 through 361		•	350 through 361
92	3 through 131	•	3 through 131	132		591 through 605
93	33 through 185	33 through 80	81 through 185	186	570 through 575	586 through 591
94	184 through 915	184 through 237	238 through 915	916	1119 through 1124	1139 through 1150
95	58 through 1116	58 through 159	160 through 1116	1117	1486 through 1491	1504 through 1513
96	327 through 417	·	327 through 417		•	404 through 417
97	63 through 398	63 through 206	207 through 398	399		•
98	2 through 163	·	2 through 163	164	488 through 493	511 through 522
99	13 through 465	13 through 75	76 through 465	466	•	
100	20 through 703	20 through 94	95 through 703	704	1000 through 1005	1023 through 1041
101	103 through 294	103 through 243	244 through 294	295	•	•
102	81 through 518	81 through 173	174 through 518	519	•	
103	66 through 326	•	66 through 326	327	1066 through 1071	1087 through 1098
104	170 through 289	170 through 250	251 through 289	290	•	•
105	36 through 497	•	36 through 497	498	650 through 655	663 through 685
106	18 through 320	•	18 through 320	321	539 through 544	542 through 554
107	71 through 1438	71 through 136	137 through 1438	1439	1644 through 1649	1665 through 1678
108	25 through 318	25 through 75	76 through 318	319	452 through 457	482 through 494
109	84 through 332	84 through 170	171 through 332	333	•	702 through 714
110	32 through 718	32 through 100	101 through 718	719	770 through 775	793 through 805
111	26 through 481	26 through 88	89 through 481	482	755 through 760	775 through 787
112	26 through 562	26 through 187	188 through 562	563	•	·
113	4 through 810	4 through 279	280 through 810	811	858 through 863	881 through 893
114	55 through 459	55 through 120	121 through 459	460	1444 through 1449	1462 through 1475
115	48 through 248	48 through 161	162 through 248	249	283 through 288	308 through 321
116	25 through 399	25 through 186	187 through 399	400	• .	
117	10 through 1137	10 through 72	73 through 1137	1138	1144 through 1149	1162 through 1173
118	72 through 704	72 through 161	162 through 704	705	772 through 777	
119	44 through 505	44 through 223	224 through 505	506		
120	25 through 393	25 through 150	151 through 393	394	734 through 739	757 through 770

-111-

CON	T. TABLE IV					·
121	58 through 1095	58 through 114	115 through 1095	1096	•	1202 through 1213
122	31 through 660	31 through 90	91. through 660	661	1288 through 1293	1307 through 1318
123	31 through 582	31 through 90	91 through 582	583	816 through 821	840 through 853
124	15 through 695	15 through 80	81 through 695	696	795 through 800	814 through 826
125	74 through 295	74 through 196	197 through 295	296	545 through 550	561 through 571
126	440 through 659	·	440 through 659	·	601 through 606	•
127	38 through 283	38 through 85	86 through 283	284	257 through 262	•
128	121 through 477	121 through 288	289 through 477		•	
129	2 through 163	•	2 through 163	164	292 through 297	310 through 323
130	46 through 675	46 through 87	88 through 675	676	1364 through 1369	1383 through 1392
131	62.through 385	•	62 through 385	386	974 through 979	987 through 999
132	422 through 550	422 through 475	476 through 550	551	•	714 through 725
133	124 through 231	•	124 through 231	232	•	387 through 400
134	131 through 1053	131 through 169	170 through 1053	•	1019 through 1024	·
135	86 through 403	86 through 181	182 through 403	404	1097 through 1102	1117 through 1128
136	37 through 162	37 through 93	94 through 162	163	224 through 229 .	243 through 254
137	31 through 381	31 through 90	91 through 381	382	•	875 through 886
138	46 through 579	46 through 156	157 through 579	580	•	•
139	92 through 471	92 through 172	173 through 471		454 through 459	458 through 471
140	154 through 675	154 through 498	499 through 675	676	819 through 824	338 through 849
242	18 through 173	18 through 77	78 through 173	174	864 through 869	882 through 893
243	17 through 595	17 through 85	86 through 595	596	820 through 825	840 through 851
244	89 through 334	89 through 130	131 through 334	335	462 through 467	484 through 495
245	21 through 614	21 through 83	84 through 614	615	849 through 854	873 through 884
246	94 through 573	94 through 258	259 through 573	574	862 through 867	886 through 897
247	74 through 397	74 through 127	128 through 397	398	472 through 477	507 through 518
248	51 through 242	51 through 116	117 through 242	243	319 through 324	339 through 350
249	111 through 191	111 through 155	156 through 191	192	965 through 970	986 through 996
250	45 through 602	45 through 107	108 through 602	603	828 through 833	850 through 860
251	24 through 560	24 through 101	102 through 560	561	563 through 568	583 through 593
252	109 through 558	109 through 273	274 through 558	559	·	1104 through 1114
253	128 through 835	128 through 220	221 through 835	836	1145 through 1150	1170 through 1181
254	59 through 505	59 through 358	359 through 505	506	1042 through 1047	1062 through 1073
255	1 through 207	1 through 147	148 through 207	208	784 through 789	807 through 818
256	12 through 734	12 through 101	102 through 734	.735	914 through 919	961 through 971
257	378 through 518	378 through 467	468 through 518	519	607 through 612	628 through 640
258	110 through 304	110 through 193	194 through 304	305	708 through 713	732 through 743
259	201 through 419	201 through 272	273 through 419	420	601 through 606	627 through 637
260	123 through 302	123 through 176	177 through 302	303	1279 through 1284	1301 through 1312
261	98 through 673	98 through 376	377 through 673	674	•	1025 through 1035
262	17 through 463	17 through 232	233 through 463	464	657 through 662	684 through 696
263	263 through 481	263 through 322	323 through 481	482	·	858 through 868

265 186 through 431 186 through 269 261 through 431 432	CONT	. TABLE IV					
268 1279 through 473 278 through 530 383 through 473 474 944 through 949 970 through 981 1676 172 through 644 12 through 92 93 through 644 645 1002 through 1007 1020 through 1031 268 91 through 489 11 through 303 331 through 489 480 . 1271 through 1281 1771 through 1281 1780 through 327 70 through 147 148 through 327 328 1741 through 1746 1763 through 1774 1770 12 through 487 12 through 104 105 through 387 384 809 through 614 632 through 667 1871 1871 1871 1871 1871 1871 1871	264	42 through 299	42 through 101	102 through 299	300	•	762 through 775
267 12 through 644 12 through 92 93 through 644 645 1002 through 1007 1020 through 1031 1268 91 through 459 91 through 330 331 through 459 460 . 1271 through 1281 1270 through 1281 770 through 1281 1270 12 through 327 70 through 147 148 through 327 328 1741 through 1746 1751 through 1774 1775 1776 1776 1776 1776 1776 1776 1776	265	198 through 431	198 through 260	261 through 431	432	l	
288 91 through 459 91 through 330 331 through 459 460	266	279 through 473	279 through 362	363 through 473	474		
288 70 through 327 70 through 147 148 through 927 328 1741 through 1774 1763 through 967 270 12 through 349 12 through 104 105 through 497 498 393 through 940 955 through 967 271 90 through 383 30 through 200 201 through 383 384 609 through 614 632 through 649 372 373 through 376 377 through 541 332 through 376 377 through 545 542 739 through 644 652 through 649 273 343 through 222 43 through 376 377 through 541 542 739 through 744 761 through 773 273 43 through 222 43 through 376 181 through 222 223 530 through 535 555 through 568 181 through 231 232 through 384 232 through 300 301 through 384 385 650 through 655 662 through 673 275 124 through 463 224 through 379 380 through 427 428 606 through 655 662 through 673 182 through 463 294 through 379 380 through 483 464 762 through 671 162 through 389 398 through 671 672 805 through 810 830 sthrough 872 278 83 through 632 63 through 380 398 through 671 672 805 through 813 829 through 829 21 through 362 21 through 363 22 through 363 308 through 673 633 808 through 813 829 through 440 280 21 through 362 21 through 364 345 through 362 363 821 through 826 838 through 849 281 21 through 362 21 through 364 345 through 503 504 1305 through 131 1300 through 134 132 through 829 21 through 201 1 through 63 64 through 201 202 837 through 842 660 through 671 182 through 672 805 through 849 849 849 849 849 849 849 849 849 849	267	12 through 644	12 through 92	93 through 644	645	1002 through 1007	
270 12 through 497 12 through 194 105 through 497 498 935 through 940 955 through 967 270 300 through 497 320 through 497 332 through 578 377 through 541 542 739 through 744 761 through 773 761 through 773 332 through 541 332 through 376 377 through 541 542 739 through 744 761 through 773 762 through 773 763 through 774 761 through 773 763 through 774 761 through 773 763 through 774 761 through 774 761 through 774 761 through 774 761 through 775 762 through 430 761 through 231 232 433 through 424 244 through 435 555 through 566 762 763 through 434 763 through 455 762 through 384 365 765 through 455 762 through 457 762 through 457 762 through 457 762 through 457 762 through 673 762 through 673 762 through 673 762 through 673 762 through 674 762 through 772 763 through 671 762 through 774 762 through 775 762 through 775 763 through 775 775 through 775 775 through 775 775 through 775 777 through 775	268	91 through 459	91 through 330	331 through 459	460	•	1271 through 1281
271 90	269	70 through 327	70 through 147	148 through 327	328		
272 32 through 541 332 through 578 377 through 541 542 739 through 744 761 through 773 273 43 through 541 332 through 578 377 through 541 542 739 through 744 761 through 773 274 115 through 222 43 through 422 43 through 424 445 through 455 555 through 455 275 232 through 344 232 through 300 301 through 427 448 through 427 445 through 455 662 through 655 662 through 655 276 143 through 427 143 through 286 287 through 427 428 606 through 611 628 through 639 277 284 through 427 143 through 286 287 through 483 398 through 483 464 - 762 through 639 278 162 through 632 63 through 308 309 through 671 672 805 through 810 830 through 840 281 21 through 632 63 through 308 309 through 671 672 805 through 840 281 21 through 302 21 through 344 345 through 503 504 1305 through 1310 1330 through 134 <t< td=""><td>270</td><td>12 through 497</td><td>12 through 104</td><td>105 through 497</td><td>498</td><td>935 through 940</td><td>955 through 967</td></t<>	270	12 through 497	12 through 104	105 through 497	498	935 through 940	955 through 967
273 43 through 222 43 through 177 178 through 222 223 530 through 535 555 through 586 274 115 through 231 115 through 180 181 through 231 232 419 through 424 445 through 455 275 232 through 384 232 through 300 301 through 384 385 650 through 655 662 through 673 276 143 through 427 143 through 286 287 through 427 428 606 through 611 628 through 639 277 284 through 631 162 through 398 399 through 643 464 762 through 671 162 through 398 399 through 643 633 805 through 631 829 through 840 278 83 through 632 63 through 308 309 through 632 633 808 through 813 829 through 840 280 21 through 302 21 through 302 303 through 303 303 through 322 21 through 503 21 through 344 345 through 503 504 1305 through 1310 1330 through 1341 282 1 through 503 21 through 63 64 through 201 202 637 through 642 660 through 671 182 through 344 345 through 503 504 1305 through 1310 1330 through 1341 284 89 through 503 69 through 125 126 through 201 1566 through 571 1587 through 1054 89 through 384 89 through 201 1 201 through 201 1 202 505 through 203 505 through 507 1587 through 1597 1 1587 through 1059 286 90 through 344 90 through 140 141 through 344 345 500 through 505 515 through 505 516 through 505 500 through 505 516 through 507 500 through 505 516 through 507 500 through 50	271	90 through 383	90 through 200	201 through 383	384	609 through 614	632 through 643
274 115 through 231 115 through 380 181 through 231 232 419 through 424 445 through 455 275 232 through 384 232 through 300 301 through 384 385 650 through 655 662 through 673 276 143 through 427 143 through 427 143 through 427 428 606 through 611 628 through 639 277 284 through 463 284 through 463 464 - 762 through 772 278 162 through 671 162 through 389 399 through 671 672 805 through 810 830 through 840 279 63 through 632 63 through 384 309 through 632 633 808 through 813 829 through 840 280 21 through 362 21 through 3840 309 through 632 633 808 through 813 829 through 840 281 21 through 636 21 through 201 201 through 840 303 through 840 303 through 840 282 1 through 503 21 through 364 64 through 344 36 through 364 660 through 361 1305 through 364 660 through 361 1305 through 364 660 th	272	332 through 541	332 through 376	377 through 541	542	739 through 744	761 through 773
275 232 through 384 232 through 300 301 through 427 428 650 through 651 662 through 673 276 143 through 427 143 through 265 287 through 427 428 606 through 611 628 through 639 277 284 through 463 294 through 379 380 through 463 484 762 through 611 830 through 840 278 162 through 671 162 through 308 399 through 671 672 805 through 810 830 through 840 280 21 through 362 21 through 200 201 through 362 383 808 through 813 829 through 849 281 21 through 362 21 through 344 345 through 362 363 through 364 37 through 826 838 through 849 281 21 through 363 41 through 363 64 through 201 202 537 through 841 135 through 134 135 through 1034 1035 through 1310 1330 through 134 135 through 1034 1035 through 1571 1587 through 1571 1587 through 1597 284 69 through 263 69 through 263 264 1173 through 178 1196 through 1597 285 115 through 285 115 through 240 205 through 341 345 </td <td>273</td> <td>43 through 222</td> <td>43 through 177</td> <td>178 through 222</td> <td>223</td> <td>530 through 535</td> <td>555 through 566</td>	273	43 through 222	43 through 177	178 through 222	223	530 through 535	555 through 566
276	274	115 through 231	115 through 180	181 through 231	232	419 through 424	445 through 455
277	275	232 through 384	232 through 300	301 through 384	385	650 through 655	662 through 673
278	276	143 through 427	143 through 286	287 through 427	428	606 through 611	628 through 639
279 63 through 632 63 through 308 309 through 632 633 808 through 813 829 through 840	277	284 through 463	284 through 379	380 through 463	464	·	762 through 772
280 21 through 362 21 through 200 201 through 362 363 821 through 826 838 through 849	278	162 through 671	162 through 398	399 through 671	672	805 through 810	830 through 840
21 1 1 1 1 1 1 1 1 1	279	63 through 632	63 through 308	309 through 632	633	808 through 813	829 through 840
281 21 through 503 21 through 344 345 through 503 504 1305 through 1310 1330 through 1341 282 1 through 201 1 through 63 64 through 201 202 637 through 642 660 through 671 283 39 through 1034 39 through 134 135 through 1034 1035 1566 through 1571 1587 through 1597 284 69 through 263 69 through 125 126 through 263 264 1173 through 1178 1196 through 1205 285 115 through 285 115 through 204 205 through 285 286 505 through 510 525 through 536 286 90 through 344 90 through 140 141 through 344 345 500 through 505 515 through 527 287 57 through 311 57 through 107 108 through 311 312 467 through 472 482 through 483 288 96 through 302 96 through 182 183 through 302 303 - 501 through 484 290 210 through 332 210 through 329 300 through 332 333 594 through 596 613 through 614 291 <td></td> <td>21 through 362</td> <td>21 through 200</td> <td>201 through 362</td> <td>363</td> <td>821 through 826</td> <td>838 through 849</td>		21 through 362	21 through 200	201 through 362	363	821 through 826	838 through 849
283 39 through 1034 39 through 134 135 through 1034 135 through 263 69 through 1571 1587 through 1597 284 69 through 263 69 through 265 126 through 263 264 1173 through 1178 1196 through 1205 285 115 through 285 115 through 204 205 through 285 286 505 through 510 525 through 536 286 30 through 344 90 through 140 141 through 344 345 500 through 505 515 through 527 287 57 through 311 57 through 107 108 through 311 312 467 through 472 482 through 493 288 96 through 302 96 through 182 183 through 302 303 501 through 472 482 through 493 289 161 through 328 329 through 302 303 594 through 576 579 through 811 290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 381 362 650 through 655 673 through 684 292	281		21 through 344	345 through 503	504	1305 through 1310	1330 through 1341
284 69 through 263 69 through 125 126 through 263 264 1173 through 1178 1196 through 1205 285 115 through 285 115 through 204 205 through 285 286 505 through 510 525 through 536 286 90 through 344 90 through 140 141 through 344 345 500 through 505 515 through 527 287 57 through 311 57 through 107 108 through 311 312 467 through 472 482 through 493 288 96 through 302 96 through 182 183 through 526 527 799 through 514 289 161 through 526 161 through 328 329 through 526 527 799 through 811 290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 576 154 through 360 361 through 576 577 737 through 782 801 through 812 294 154 through 576 154 through 360 361 through 576 577 737 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 298 66 through 497 66 through 253 254 through 897 898 1017 through 1022 1044 through 1054 298 66 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 383 126 through 69 7 through 497 498 594 through 799 618 through 629 299 49 through 534 49 through 69 7 through 534 535 593 through 598 612 through 623 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 103 104 through 415 416 540 through 513 528 through 539 302 56 through 527 21 through 99 96 through 527 528 921 through 926 953 through 539 304 21 through 527 21 through 59 96 through 527 528 921 through 926 953 through 539	282	1 through 201	1 through 63	64 through 201	202	637 through 642	660 through 671
285 115 through 285 115 through 294 205 through 285 286 505 through 510 525 through 536 286 90 through 344 90 through 140 141 through 344 345 500 through 505 515 through 527 287 57 through 311 57 through 107 108 through 311 312 467 through 472 482 through 493 288 96 through 302 96 through 182 183 through 302 303 501 through 514 289 161 through 526 161 through 289 329 through 526 527 799 through 811 290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 682 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 360 361 through 576 577 737 through 782 801 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 497 498 594 through 599 618 through 629 299 49 through 491 49 through 96 97 through 491 492 593 through 599 618 through 623 300 49 through 534 49 through 96 97 through 534 593 through 599 618 through 623 301 86 through 588 56 through 145 146 through 105 101 through 589 593 through 599 601 through 623 301 86 through 588 56 through 145 146 through 105 101 through 596 584 through 598 612 through 623 301 86 through 588 56 through 145 146 through 497 498 594 through 599 601 through 623 301 86 through 588 56 through 100 101 through 588 593 through 589 601 through 571 302 56 through 527 21 through 100 101 through 527 528 921 through 589 601 through 513 528 through 539 304 21 through 527 21 through 599 61 through 527 528 921 through 596 953 through 583 583 through 589 601 through 513 528 through 589 601 through 513 528 through 589 601 through 513 528 through 589 601 through 503 304 21 through 527 21 through 599 96 through 527 528 921 through 596 953 through 583 304 21 throu	283	39 through 1034	39 through 134	135 through 1034	1035	1566 through 1571	1587 through 1597
286 90 through 344 90 through 140 141 through 344 345 500 through 505 515 through 527 287 57 through 311 57 through 107 108 through 311 312 467 through 472 482 through 493 288 96 through 302 96 through 182 183 through 302 303 501 through 514 289 161 through 526 161 through 328 329 through 526 527 799 through 811 290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 897 154 through 360 361 through 897 898 1017 through 742 763 through 754 295 154 through 897 154 throug	284	69 through 263	69 through 125	126 through 263	264	1173 through 1178	1196 through 1205
287 57 through 311 57 through 107 108 through 311 312 487 through 472 482 through 493 288 96 through 302 96 through 182 183 through 302 303 501 through 514 289 161 through 526 161 through 328 329 through 526 527 799 through 811 290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 360 361 through 576 577 737 through 742 763 through 775 295 154 through 387 154 through 897 898 1017 through 742 763 through 754 296 146 through 292 146 through 253 254 thro	285	115 through 285	115 through 204	205 through 285	286	505 through 510	525 through 536
288 96 through 302 96 through 182 183 through 302 303 501 through 514 289 161 through 526 161 through 328 329 through 526 527 799 through 811 290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 360 361 through 576 577 737 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 383 126 through 253 254 through 383 384 726 through 731 743 through 743 through 754 298 66 through 497	286	90 through 344	90 through 140	141 through 344	345	500 through 505	515 through 527
288 161 through 526 161 through 328 329 through 526 527 799 through 811 290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 576 577 737 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 293 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66	287	57 through 311	57 through 107	108 through 311	312	467 through 472	482 through 493
290 210 through 332 210 through 299 300 through 332 333 594 through 599 613 through 625 291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 360 361 through 897 898 1017 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49	288	96 through 302	96 through 182	183 through 302	303	•	501 through 514
291 212 through 361 212 through 319 320 through 361 362 650 through 655 673 through 684 292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 360 361 through 576 577 737 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 300 49 through 534 49 through 145 146 through 415 416<	289	161 through 526	161 through 328	329 through 526	527	·	799 through 811
292 75 through 482 75 through 128 129 through 482 483 595 through 600 618 through 627 293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 360 361 through 897 577 737 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 534 535 593 through 737 750 through 623 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 145 146 through 268 560 through 589 601 thr	290	210 through 332	210 through 299	300 through 332	333	594 through 599	613 through 625
293 50 through 631 50 through 244 245 through 631 632 777 through 782 801 through 812 294 154 through 576 154 through 360 361 through 576 577 737 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 411 412 732 through 737 750 through 763 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 415 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 328 329 508 through 513 <td>291</td> <td>212 through 361</td> <td>212 through 319</td> <td>320 through 361</td> <td>362</td> <td>650 through 655</td> <td>673 through 684</td>	291	212 through 361	212 through 319	320 through 361	362	650 through 655	673 through 684
294 154 through 576 154 through 360 361 through 576 577 737 through 742 763 through 775 295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 239 240 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 411 412 732 through 737 750 through 763 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 415 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 95 96 through 527 528	292	75 through 482	75 through 128	129 through 482	483	595 through 600	618 through 627
295 154 through 897 154 through 360 361 through 897 898 1017 through 1022 1044 through 1054 296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 411 412 732 through 737 750 through 763 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 415 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 527 21 through 527 528 921 through 926	293	50 through 631	50 through 244	245 through 631	632	777 through 782	801 through 812
296 146 through 292 146 through 253 254 through 292 293 395 through 400 433 through 444 297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 411 412 732 through 737 750 through 763 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 415 146 through 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	294	154 through 576	154 through 360	361 through 576	577	737 through 742	763 through 775
297 126 through 383 126 through 167 168 through 383 384 726 through 731 743 through 754 298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 411 412 732 through 737 750 through 763 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 145 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 527 528 921 through 926 953 through 963	295	154 through 897	154 through 360	361 through 897	898	1017 through 1022	1044 through 1054
298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 411 412 732 through 737 750 through 763 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 415 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	296	146 through 292	146 through 253	254 through 292	293	395 through 400	433 through 444
298 66 through 497 66 through 497 498 594 through 599 618 through 629 299 49 through 411 49 through 96 97 through 411 412 732 through 737 750 through 763 300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 415 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	297	126 through 383	126 through 167	168 through 383 .	384	726 through 731	743 through 754
300 49 through 534 49 through 96 97 through 534 535 593 through 598 612 through 623 301 86 through 415 86 through 145 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963			66 through 239	240 through 497	498	594 through 599	618 through 629
301 86 through 415 86 through 145 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	299	49 through 411	49 through 96	97 through 411	412	732 through 737	750 through 763
301 86 through 415 86 through 145 146 through 415 416 540 through 545 560 through 571 302 56 through 268 56 through 100 101 through 268 269 584 through 589 601 through 612 303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	300	49 through 534	49 through 96	97 through 534	535	593 through 598	612 through 623
303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	301		86 through 145	146 through 415	416	540 through 545	560 through 571
303 32 through 328 32 through 103 104 through 328 329 508 through 513 528 through 539 304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	302	56 through 268	56 through 100	101 through 268	269	584 through 589	601 through 612
304 21 through 527 21 through 95 96 through 527 528 921 through 926 953 through 963	303		32 through 103	104 through 328	329	508 through 513	528 through 539
CC0 41 501		21 through 527	21 through 95	96 through 527	528	921 through 926	953 through 963
		147 through 647	147 through 374	375 through 647	648	•	668 through 681

CONT. TABLE IV

CON	T. TABLE IV					•
306	262 through 471	262 through 306	307 through 471	472	663 through 668	682 through 693
307	74 through 1216	74 through 172	173 through 1216	1217	1627 through 1632	1640 through 1652
308	48 through 164	48 through 89	90 through 164	165	482 through 487	505 through 517
309	185 through 334	185 through 295	296 through 334	335	355 through 360	392 through 405
310	195 through 347	195 through 272	273 through 347	348	1037 through 1042	1071 through 1082
311	90 through 815	90 through 179	180 through 815	816	883 through 888	905 through 916
312	52 through 513	52 through 231	232 through 513	514	553 through 558	572 through 583
313	172 through 438	172 through 354	355 through 438	439	682 through 687	685 through 697
314	148 through 366	148 through 225	226 through 366	367	770 through 775	792 through 803
315	175 through 336	175 through 276	277 through 336	337	-	812 through 823
316	191 through 553	191 through 304	305 through 553	554	766 through 771	804 through 817
317	106 through 603	106 through 216	217 through 603	604	·	1102 through 1112
318	47 through 586	47 through 124	125 through 586	587	1583 through 1588	1614 through 1623
319	99 through 371	99 through 290	291 through 371	372	491 through 496	513 through 524
320	44 through 814	44 through 112	113 through 814	815	.	978 through 989
321	3 through 581	3 through 182	183 through 581	582		1006 through 1016
322	107 through 427	107 through 190	191 through 427	428	499 through 504	516 through 529
323	45 through 407	45 through 83	84 through 407	408	1008 through 1013	1032 through 1042
324	201 through 332	201 through 251	252 through 332	333		869 through 880
325	217 through 543	217 through 255	256 through 543	544	•	1206 through 1217
326	18 through 446	18 through 140	141 through 446	447	930 through 935	948 through 959
327	29 through 724	29 through 118	119 through 724	725	886 through 891	910 through 920
328	404 through 586	404 through 466	467 through 586	587	1304 through 1309	1334 through 1344
329	331 through 432	331 through 387	388 through 432	433	548 through 553	573 through 585
330	59 through 703	59 through 220	221 through 703	704	886 through 891	903 through 914
331	672 through 752	672 through 722	723 through 752	753	•	1150 through 1161
332	57 through 311	57 through 128	129 through 311	312	332 through 337	351 through 363
333	80 through 232	80 through 127	128 through 232	233	617 through 622	634 through 645
334	91 through 291	91 through 219	220 through 291	292	367 through 372	389 through 400
335	196 through 384	196 through 240	241 through 384	385	461 through 466	485 through 496
336	54 through 590	54 through 227	228 through 590	591		955 through 965
337	133 through 846	133 through 345	346 through 846	847		890 through 901
338	138 through 671	138 through 248	249 through 671	672	1319 through 1324	1338 through 1347
339	124 through 411	124 through 186	187 through 411	412	948 through 953	971 through 983
340	372 through 494	372 through 443	444 through 494	495	708 through 713	732 through 745
341	112 through 450	112 through 192	193 through 450	451	1053 through 1058	1095 through 1106
342	117 through 866	117 through 170	171 through 866	867	1159 through 1164	1178 through 1190
343	13 through 465	13 through 75	76 through 465	466	1035 through 1040	1060 through 1070
344	2 through 718	2 through 76	77 through 718	719	1170 through 1175	1203 through 1213
345	86 through 709	86 through 361	362 through 709	710	943 through 948	963 through 973
346 347	63 through 320	63 through 179	180 through 320	321	771 through 776	799 through 810

CONT. TABLE IV

LUN	I. I ABLE IV					•
348	186 through 380	186 through 233	234 through 380	381	383 through 388	396 through 409
349	69 through 458	69 through 233	234 through 458	459	564 through 569	602 through 613
350	12 through 638	12 through 263	264 through 638	639	951 through 956	975 through 985
351	282 through 389	282 through 332	333 through 389	390	1413 through 1418	1437 through 1447
352	208 through 339	208 through 294	295 through 339	340		1631 through 1641
353	69 through 557	69 through 224	225 through 557	558	849 through 854	870 through 883
354	134 through 325	134 through 274	275 through 325	326		718 through 729
355	78 through 731	78 through 227	228 through 731	732		1002 through 1013
356	46 through 693	46 through 90	91 through 693	694	937 through 942	962 through 973
357	126 through 527	126 through 182	183 through 527	528	834 through 839	856 through 867
358	66 through 320	66 through 113	114 through 320	321	490 through 495	508 through 519
359	73 through 948	73 through 159	160 through 948	949	•	1016 through 1028
360	69 through 434	69 through 236	237 through 434	435	419 through 424	441 through 452
361	628 through 804	628 through 711	712 through 804	805		864 through 875
362	70 through 366	70 through 108	109 through 366	367	496 through 501	521 through 531
363	70 through 366	70 through 108	109 through 366	367	· · · · · · · · · · · · · · · · · · ·	1233 through 1244
364	111 through 434	111 through 185	186 through 434	435		618 through 631
365	19 through 567	19 through 63	64 through 567	568	749 through 754	771 through 781
366	19 through 312	19 through 63	64 through 312	313	896 through 901	921 through 931
367	64 through 612	64 through 234	235 through 612	613	•	839 through 849
368	39 through 458	39 through 80	81 through 458	459	613 through 618	633 through 644
369	9 through 185	9 through 50	51 through 185	186	•	906 through 918
370	14 through 316	14 through 121	122 through 316	317	442 through 447	458 through 471
371	70 through 1092	70 through 234	235 through 1092	1093	1475 through 1480	1493 through 1504
372	274 through 597	274 through 399	400 through 597	598	731 through 736	754 through 765
373	230 through 469	230 through 307	308 through 469	470	1004 through 1009	1027 through 1040
374	72 through 545	72 through 203	204 through 545	546		1151 through 1162
375	36 through 425	36 through 119	120 through 425	426	1215 through 1220	1240 through 1250
376	155 through 751	155 through 340	341 through 751	752	912 through 917	937 through 947
377	46 through 585	46 through 120	121 through 585	586	584 through 589	606 through 619

TABLE V

		I ABLE V	
ld	Full Length Polypeptide Location	Signal Peptide Location	Mature Polypeptide Location
141	-31 through 124	-31 through -1	1 through 124
142	1 through 55		1 through 55
143	-20 through 47	-20 through -1	1 through 47
144	-21 through 177	-21 through -1	1 through 177
145	-25 through 110	-25 through -1	1 through 110
146	-70 through 185	-70 through -1	1 through 185
147	-49 through 10	-49 through -1	1 through 10
148	1 through 180		1 through 180
149	-23 through 139	-23 through -1	1 through 139
150	-23 through 97	-23 through -1	1 through 97
151	1 through 7		1 through 7
152	-42 through 157	-42 through -1	
153	1 through 43		1 through 157
154	-37 through 13	-37 through -1	1 through 43
155	1 through 153		1 through 13
156	1 through 67		1 through 153
157	1 through 87		1 through 67
158	-85 through 165	-85 through -1	1 through 87
159	1 through 24	-03 through -1	1 through 165
160	1 through 228		1 through 24
161	-20 through 66	-20 through -1	1 through 228
162	1 through 44	-20 tillough - 1	1 through 66
163	-58 through 256	-58 through -1	1 through 44
164	-80 through 9	-80 through -1	1 through 256
165	-15 through 83	-15 through -1	1 through 9
166	-36 through 56	-36 through -1	1 through 83
167	-16 through 335	-16 through -1	1 through 56
168	-47 through 91	-47 through -1	1 through 335
169	-73 through 28	-73 through -1	1 through 91
170	-68 through 184	-68 through -1	1 through 28
171	-68 through 282	-68 through -1	1 through 184
172	-68 through 322	-68 through -1	1 through 282
173	-82 through 108	-82 through -1	1 through 322
174	-232 through 53	-232 through -1	1 through 108
175	1 through 153	-232 tillough - 1	1 through 53
176	1 through 49		1 through 153
177	-24 through 75	-24 through -1	1 through 49
178	-37 through 58	-37 through -1	1 through 75
179	-23 through 98	-23 through -1	1 through 58
180	1 through 59	zo mrodyn - i	1 through 98
181	-14 through 72	-14 through -1	1 through 59
182	-58 through 107	-58 through -1	1 through 72
183	-35 through 45	-35 through -1	1 through 107
184	·21 through 52	-21 through -1	1 through 45
185	1 through 98	er anough -1	1 through 52
186	-21 through 91	-21 through -1	1 through 98
187	-44 through 26	-44 through -1	1 through 91
188	-13 through 79	-13 through -1	1 through 26
189	-42 through 165	-42 through -1	1 through 79 1 through 165
190	1 through 201	76 UNUUHII 1	i mromo its

NT. TABLE V			
191	-37 through 342	-37 through -1	1 through 342
192	1 through 112	· · · · · · · · · · · · · · · · · · ·	1 through 112
193	1 through 43	<u> </u>	1 through 43
194	-16 through 35	-16 through -1	1 through 35
195	-18 through 226	-18 through -1	1 through 226
196	-34 through 319	-34 through -1	1 through 319
197	1 through 30		1 through 30
198	-48 through 64	-48 through -1	1 through 64
199	1 through 54		1 through 54
200	-21 through 130	-21 through -1	1 through 130
201	-25 through 203	-25 through -1	1 through 203
202	-47 through 17	-47 through -1	1 through 17
203	-31 through 115	-31 through -1	1 through 115
204	1 through 87	•	1 through 87
205	-27 through 13	-27 through -1	1 through 13
206	1 through 154	•	1 through 154
207	1 through 101		1 through 101
208	-22 through 434	-22 through -1	1 through 434
209	-17 through 81	-17 through -1	1 through 81
210	-29 through 54	-29 through -1	1 through 54
211	-23 through 206	-23 through -1	1 through 206
212	-21 through 131	-21 through -1	1 through 131
213	-54 through 125	-54 through -1	1 through 125
214	-92 through 177	-92 through -1	1 through 177
215	-22 through 113	-22 through -1	1 through 113
216	-38 through 29	-38 through -1	1 through 29
217	-54 through 71	-54 through -1	1 through 71
218	-21 through 355	-21 through -1	1 through 355
219	-30 through 181	-30 through -1	1 through 181
220	-60 through 94	-60 through -1	1 through 94
221	-42 through 81	-42 through -1	1 through 81
222	-19 through 327	-19 through -1	1 through 327
223	-20 through 190	-20 through -1	1 through 190
224	-20 through 164	-20 through -1	1 through 164
225	-22 through 205	-22 through -1	1 through 205
226	-41 through 33	-41 through -1	1 through 33
227	1 through 73		1 through 73
228	-16 through 66	-16 through -1	1 through 66
229	-56 through 63	-56 through -1	1 through 63
230	1 through 54		1 through 54
231	-14 through 196	-14 through -1	1 through 196
232	1 through 108		1 through 108
233	-18 through 25	-18 through -1	1 through 25
234	1 through 36		1 through 36
235	-13 through 294	-13 through -1	1 through 294
236	-32 through 74	-32 through -1	1 through 74
237	-19 through 23	-19 through -1	1 through 23
238	-20 through 97	-20 through -1	1 through 97
239	-37 through 141	-37 through -1	1 through 141
240	-27 through 99	-27 through -1	1 through 99
241	-115 through 59	-115 through -1	
			1 through 59
378	-20 through 32	•2U through .1	1 through 22
378 379	-20 through 32 -23 through 170	-20 through -1 -23 through -1	1 through 32 1 through 170

		~ .	-	_	11
CO.	M I	ΙΔ	ĸı	-	w
L U	IW I .	TA	טע		•

NT. TABLE V	-21 through 177	-21 through -1	1 through 177
382	-55 through 105	-55 through -1	1 through 105
383	-18 through 90	-18 through -1	1 through 90
384	-22 through 42	-22 through -1	1 through 42
385	-15 through 12	-15 through -1	1 through 12
386	-21 through 165	-21 through -1	1 through 165
387	-26 through 153	-26 through -1	1 through 153
388	-55 through 95	-55 through -1	1 through 95
389	-31 through 205	-31 through -1	1 through 205
390	-100 through 49	-100 through -1	1 through 49
391	-49 through 20	-49 through ⋅1	1 through 20
392	-30 through 211	-30 through -1	1 through 211
393	-30 through 17	-30 through -1	1 through 17
394	-28 through 37	-28 through -1	1 through 37
395	-24 through 49	-24 through -1	1 through 49
396	-18 through 42	-18 through -1	1 through 42
	-93 through 99	-93 through -1	1 through 99
397	-72 through 77	-72 through -1	1 through 77
398	-20 through 53	-20 through -1	1 through 53
399		-20 through -1	1 through 66
400	-20 through 66 -21 through 57	-21 through -1	1 through 57
401		-28 through -1	1 through 37
402	-28 through 37	-27 through -1	1 through 184
403	-27 through 184	-80 through -1	1 through 43
404	-80 through 43	-26 through -1	1 through 60
405	-26 through 60	-31 through -1	1 through 131
406	-31 through 131	-37 through -1	1 through 61
407	-37 through 61	-15 through -1	1 through 55
408	-15 through 55	-45 through -1	1 through 15
409	-45 through 15	-22 through -1	1 through 17
410	-22 through 17		1 through 28
411	-23 through 28	-23 through -1 -48 through -1	1 through 47
412	-48 through 47		1 through 28
413	-32 through 28	-32 through -1	1 through 91
414	-79 through 91	-79 through -1	1 through 108
415	-82 through 108	-82 through -1	1 through 54
416	-60 through 54	-60 through -1	1 through 53
417	-108 through 53	-108 through -1	
418	-21 through 46	-21 through -1	1 through 46 1 through 300
419	-32 through 300	-32 through -1	1 through 46
420	-19 through 46	-19 through -1	1 through 48
422	-30 through 27	-30 through -1	
423	-17 through 68	-17 through -1	1 through 68
424	-17 through 68	-17 through -1	1 through 68
425	-29 through 40	-29 through -1	1 through 40
426	-56 through 66	-56 through -1	1 through 66
427	-30 through 11	-30 through -1	1 through 11
428	-36 through 14	-36 through -1	1 through 14
429	-18 through 118	-18 through -1	1 through 118
430	-65 through 129	-65 through -1	1 through 129
431	-69 through 72	-69 through -1	1 through 72
432	-69 through 179	-69 through -1	1 through 179
433	-36 through 13	-36 through -1	1 through 13
434	-14 through 72	-14 through -1	1 through 72
435	-58 through 86	-58 through -1	1 through 86

CONT. TABLE V

ONT. TABLE V			
436	-16 through 105	-16 through -1	1 through 105
437	-16 through 146	-16 through -1	1 through 146
438	-20 through 90	-20 through -1	1 through 90
439	-15 through 56	-15 through -1	1 through 56
440	-24 through 75	-24 through -1	1 through 75
441	-25 through 144	-25 through -1	1 through 144
442	-76 through 91	-76 through -1	1 through 91
443	-15 through 55	-15 through -1	1 through 55
444	-33 through 348	-33 through -1	1 through 348
445	-14 through 25	-14 through -1	1 through 25
446	-37 through 13	-37 through -1	1 through 13
447	-26 through 25	-26 through -1	1 through 25
448	-30 through 212	-30 through -1	1 through 212
449	-60 through 94	-60 through -1	1 through 94
450	-61 through 28	-61 through -1	1 through 28
451	-26 through 47	-26 through -1	1 through 47
452	-34 through 20	-34 through -1	1 through 20
453	-38 through 83	-38 through -1	1 through 83
454	-37 through 129	-37 through -1	1 through 129
455	-26 through 154	-26 through -1	1 through 154
456	-64 through 27	-64 through -1	1 through 27
457	-23 through 234	-23 through -1	1 through 234
458	-60 through 133	-60 through -1	1 through 133
459	-28 through 79	-28 through -1	1 through 79
460	-13 through 108	-13 through -1	1 through 108
461	-17 through 27	-17 through -1	1 through 27
462	-13 through 96	-13 through -1	1 through 96
463	-41 through 102	-41 through -1	1 through 102
464	-30 through 202	-30 through -1	1 through 202
465	-21 through 40	-21 through -1	1 through 40
466	-19 through 15	-19 through -1	1 through 15
467	-54 through 161	-54 through -1	1 through 161
468	-17 through 10	-17 through -1	1 through 10
469	-24 through 61	-24 through -1	1 through 61
470	-16 through 35	-16 through -1	1 through 35
471	-43 through 24	-43 through -1	1 through 24
472	-15 through 48	-15 through -1	1 through 48
473	-58 through 121	-58 through -1	1 through 121
474	-71 through 167	-71 through -1	1 through 167
475	-37 through 141	-37 through -1	1 through 141
476	-21 through 75	-21 through -1	1 through 75
477	-24 through 17	-24 through -1	1 through 17
478	-27 through 86	-27 through -1	1 through 86
479	-18 through 232	-18 through -1	1 through 232
480	-21 through 130	-21 through -1	1 through 130
481	-25 through 214	-25 through -1	1 through 214
482	-92 through 116	-92 through -1	1 through 116
483	-39 through 47	-39 through -1	1 through 47
484	-27 through 13	-27 through -1	1 through 13
485	-16 through 49	-16 through -1	1 through 49
486	-55 through 75	-55 through -1	1 through 75
487	-84 through 125	-84 through -1	1 through 125
488	-17 through 19	-17 through -1	1 through 19
489	-29 through 15	-29 through -1	. 1 through 15

490	-52 through 111	-52 through -1	1 through 111
491	-47 through 17	-47 through -1	1 through 17
492	-50 through 168	-50 through -1	1 through 168
493	-15 through 201	-15 through -1	1 through 201
494	-19 through 115	-19 through -1	1 through 115
495	-16 through 69	-16 through -1	1 through 69
496	-29 through 263	-29 through -1	1 through 263
497	-56 through 66	-56 through -1	1 through 66
498	-28 through 31	-28 through -1	1 through 31
499	-13 through 86	-13 through -1	1 through 86
500	-13 through 86	-13 through -1	1 through 86
501	-25 through 83	-25 through -1	1 through 83
502	-15 through 168	-15 through -1	1 through 168
503	-15 through 83	-15 through -1	1 through 83
504	-57 through 126	-57 through -1	1 through 126
505	-14 through 126	-14 through -1	1 through 126
506	-14 through 45	-14 through -1	1 through 45
507	-36 through 65	-36 through -1	1 through 65
508	-55 through 286	-55 through -1	1 through 286
509	-42 through 66	-42 through -1	1 through 66
510	-26 through 54	-26 through -1	1 through 54
511	-44 through 114	-44 through -1	1 through 114
512	-28 through 102	-28 through -1	1 through 102
513	-62 through 137	-62 through -1	1 through 137
514	-25 through 155	-25 through -1	1 through 155

-120-

TABLE VI

ld	Collection refs	Deposit Name
40	ATCC # 98921	SignalTag 121-144
41	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
42	ATCC # 98921	SignalTag 121-144
43	ATCC # 98920	SignalTag 67-90
44	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
45	ATCC # 98920	SignalTag 67-90
46	ATCC # 98923	SignalTag 44-66
47	ATCC # 98920	SignalTag 67-90
48	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
49	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
50	ATCC # 98921	SignalTag 121-144
51	ATCC # 98921	SignalTag 121-144
52	ATCC # 98920	SignalTag 67-90
53	ATCC # 98923	SignalTag 44-66
54	ATCC # 98920	SignalTag 67-90
55	ATCC # 98920	SignalTag 67-90
56	ATCC # 98920	SignalTag 67-90
57	ATCC # 98921	SignalTag 121-144
58	ATCC # 98920	SignalTag 67-90
59	ATCC # 98920	SignalTag 67-90
60	ATCC # 98920	SignalTag 67-90
61	ATCC # 98923	SignalTag 44-66
62	ATCC # 98923	SignalTag 44-66
63	ATCC # 98923	SignalTag 44-66
64	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
65	ATCC # 98923	SignalTag 44-66
66	ATCC # 98921	SignalTag 121-144
67	ATCC # 98920	SignalTag 67-90
68	ATCC # 98920	SignalTag 67-90
69	ATCC # 98921	SignalTag 121-144
70	ATCC # 98921	SignalTag 121-144
71	ATCC # 98921	SignalTag 121-144
72	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
'3	ATCC # 98923	SignalTag 44-66

74	ATCC # 98923	SignalTag 44-66
75	ATCC # 98920	SignalTag 67-90
76	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
77	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
78	ATCC # 98921	SignalTag 121-144
79	ATCC # 98923	SignalTag 44-66
80	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
81	ATCC # 98921	SignalTag 121-144
82	ATCC # 98920	SignalTag 67-90
83	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
84	ATCC # 98923	SignalTag 44-66
85	ATCC # 98923	SignalTag 44-66
86	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
87	ATCC # 98923	SignalTag 44-66
88	ATCC # 98923	SignalTag 44-66
89	ATCC # 98923	SignalTag 44-66
90	ATCC # 98923	SignalTag 44-66
91	ATCC # 98923	SignalTag 44-66
92	ATCC # 98920	SignalTag 67-90
93	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
94	ATCC # 98923	SignalTag 44-66
95	ATCC # 98923	SignalTag 44-66
96	ATCC # 98920	SignalTag 67-90
97	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
98	ATCC # 98921	SignalTag 121-144
99	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
100	ATCC # 98921	SignalTag 121-144
101	ATCC # 98920	SignalTag 67-90
102	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
103	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
104	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
105	ATCC # 98921	SignalTag 121-144
106	ATCC # 98920	SignalTag 67-90
107	ATCC # 98920	SignalTag 67-90
108	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
109	ATCC # 98923	SignalTag 44-66
10	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120

111	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
112	ATCC # 98920	SignalTag 67-90
113	ATCC # 98920	SignalTag 67-90
114	ATCC # 98923	SignalTag 44-66
115	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
116	ATCC # 98920	SignalTag 67-90
117	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
118	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
119	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
120	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
121	ATCC # 98923	SignalTag 44-66
122	ATCC # 98920	SignalTag 67-90
123	ATCC # 98920	SignalTag 67-90
124	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
125	ECACC # 98121506	SignalTag 11121998
126	ECACC # 98121506	SignalTag 11121998
127	ECACC # 98121506	SignalTag 11121998
128	ECACC # 98121506	SignalTag 11121998
129	ECACC # 98121506	SignalTag 11121998
130	ECACC # 98121506	SignalTag 11121998
131	ECACC # 98121506	SignalTag 11121998
132	ECACC # 98121506	SignalTag 11121998
133	ECACC # 98121506	SignalTag 11121998
134	ECACC # 98121506	SignalTag 11121998
135	ECACC # 98121506	SignalTag 11121998
136	ECACC # 98121506	SignalTag 11121998
137	ECACC # 98121506	SignalTag 11121998
138	ECACC # 98121506	SignalTag 11121998
139	ECACC # 98121506	SignalTag 11121998
140	ECACC # 98121506	SignalTag 11121998

-123-

TABLE VII

		FE AII
Internal designation number	SEQ ID NO	Type of sequence
20-5-2-C3-CL0_4	40	DNA
20-8-4-A11-CL2_6	41	DNA
21-1-4-F2-CL11_1	42	DNA
22-11-2-H9-CL1_1	43	DNA
25-7-3-D4-CL0_2	44	DNA
26-27-3-D7-CL0_1	45	DNA
26-35-4-H9-CL1_1	46	DNA
26-45-2-C4-CL2_6	47	DNA
27-1-2-B3-CL0_1	48	DNA
27-1-2-B3-CL0_2	49	DNA
27-19-3-G7-CL11_2	50	DNA
33-10-4-E2-CL13_4	51	DNA
33-10-4-H2-CL2_2	52	DNA
33-110-4-A5-CL1_1	53	DNA
33-13-1-C1-CL1_1	54	DNA
33-30-2-A6-CLO_1	55	DNA
33-35-4-F4-CL1_2	56	DNA
33-35-4-G1-CL1_2	57	DNA
33-36-3-E2-CL1_1	58	DNA
33-36-3-E2-CL1_2	59	DNA
33-36-3-F2-CL2_2	60	DNA .
33-4-2-G5-CL2_1	61	DNA
33-49-1-H4-CL1_1	62	DNA
33-66-2-B10-CL4_1	63	DNA
33-97-4-G8-CL2_2	64	DNA
33-98-4-C1-CL1_3	-65	DNA
47-14-1-C3-CL0_5	66	DNA
47-15-1-E11-CLO_1	67	DNA
47-15-1-H8-CLO_2	68	DNA
48-1-1-H7-CLO_1	69	DNA
48-1-1-H7-CLO_4	70	DNA
48-1-1-H7-CLO_5	71	DNA
48-3-1-H9-CLO_6	72	DNA
48-54-1-G9-CL2_1	73	DNA

48-54-1-G9-CL3_1	74	DNA
48-7-4-H2-CL2_2	75	DNA
51-11-3-D5-CL1_3	76	DNA
51-11-3-G9-CL0_1	77	DNA
51-15-4-A12-CL11_3	78	DNA
51-17-4-A4-CL3_1	79	DNA
51-2-3-F10-CL1_5	80	DNA
51-2-4-F5-CL11_2	81	DNA
51-27-4-F2-CL0_2	82	DNA .
51-34-3-FB-CL0_2	83	DNA
57-1-4-E2-CL1_2	84	DNA
57-19-2-G8-CL2_1	85	DNA
57-27-3-G10-CL2_2	86	DNA
58-33-3-B4-CL1_2	87	DNA
58-34-3-C9-CL1_2	88	DNA
58-4-4-G2-CL2_1	89	DNA
58-48-1-G3-CL2_4	90	DNA
58-6-1-H4-CL1_1	91	DNA
60-12-1-E11-CL1_2	92	DNA
65-4-4-H3-CL1_1	93	DNA
74-5-1-E4-CL1_2	94	DNA
76-13-3-A9-CL1_2	95	DNA
76-16-1-D6-CL1_1	96	DNA
76-28-3-A12-CL1_5	97	DNA
76-42-2-F3-CL0_1	98	DNA
77-16-4-G3-CL1_3	99	DNA
77-39-4-H4-CL11_4	100	DNA
78-24-3-H4-CL2_1	101	DNA
78-27-3-D1-CL1_6	102	DNA
78-28-3-D2-CLO_2	103	DNA
78-7-1-G5-CL2_6	104	DNA
84-3-1-G10-CL11_6	105	DNA
58-48-4-E2-CLO_1	106	DNA
23-12-2-G6-CL1_2	107	DNA
25-8-4-B12-CL0_5	108	DNA
26-44-3-C5-CL2_1	109	DNA
27-1-2-B3-CLO_3	110	DNA
	<u></u>	

30-12-3-G5-CLO_1	111	DNA
33-106-2-F10-CL1_3	112	DNA
33-28-4-D1-CLO_1	113	DNA
33-31-3-C8-CL2_1	114	DNA
48-24-1-D2-CL3_2	115	DNA
48-46-4-A11-CL1_4	116	DNA
51-1-4-C1-CLO_2	117	DNA
51-39-3-H2-CL1_2	118	DNA
51-42-3-F9-CL1_1	119	DNA
51-5-3-G2-CL0_4	120	DNA
57-18-4-H5-CL2_1	121	DNA
76-23-3-G8-CL1_1	122	DNA
76-23-3-G8-CL1_3	123	DNA
78-8-3-E6-CL0_1	124	DNA
19-10-1-C2-CL1_3	125	DNA
33-11-1-B11-CL1_2	126	DNA
33-113-2-B8-CL1_2	127	DNA
33-19-1-C11-CL1_1	128	DNA
33-61-2-F6-CL0_2	129	DNA
47-4-4-C6-CL2_2	130	DNA
48-54-1-G9-CL1_1	131	DNA
51-43-3-G3-CL0_1	132	DNA
55-1-3-D11-CL0_1	133	DNA
58-14-2-D3-CL1_2	134	DNA
58-35-2-B6-CL2_3	135	DNA
76-18-1-F6-CL1_1	136	DNA
76-23-3-G8-CL2_2	137	DNA
76-30-3-B7-CL1_1	138	DNA
78-21-3-G7-CL2_1	139	DNA
58-45-4-B11-CL13_2	140	DNA
20-5-2-C3-CL0_4	141	PRT
20-8-4-A11-CL2_6	142	PRT
21-1-4-F2-CL11_1	143	PRT
22-11-2-H9-CL1_1	144	PRT
25-7-3-D4-CLO_2	145	PRT
26-27-3-D7-CLO_1	146	PRT
26-35-4-H9-CL1_1	147	PRT

26-45-2-C4-CL2_6	148	PRT
27-1-2-B3-CLO_1	149	PRT
27-1-2-B3-CLO_2	150	PRT
27-19-3-G7-CL11_2	151	PRT
33-10-4-E2-CL13_4	152	PRT
33-10-4-H2-CL2_2	153	PRT
33-110-4-A5-CL1_1	154	PRT
33-13-1-C1-CL1_1	155	PRT
33-30-2-A6-CLO_1	156	PRT
33-35-4-F4-CL1_2	157	PRT
33-35-4-G1-CL1_2	158	PRT
33-36-3-E2-CL1_1	159	PRT
33-36-3-E2-CL1_2	160	PRT
33-36-3-F2-CL2_2	161	PRT
33-4-2-G5-CL2_1	162	PRT
33-49-1-H4-CL1_1	163	PRT
33-66-2-B10-CL4_1	164	PRT
33-97-4-G8-CL2_2	165	PRT
33-98-4-C1-CL1_3	166	PRT
47-14-1-C3-CL0_5	167	PRT
47-15-1-E11-CLO_1	168	PRT
47-15-1-H8-CL0_2	169	PRT
48-1-1-H7-CLO_1	170	PRT
48-1-1-H7-CLO_4	171	PRT
48-1-1-H7-CLO_5	172	PRT
48-3-1-H9-CLO_6	173	PRT
48-54-1-G9-CL2_1	174	PRT
48-54-1-G9-CL3_1	175	PRT
48-7-4-H2-CL2_2	176	PRT
51-11-3-D5-CL1_3	177	PRT
51-11-3-G9-CLO_1	178	PRT
51-15-4-A12-CL11_3	179	PRT
51-17-4-A4-CL3_1	180	PRT
51-2-3-F10-CL1_5	181	PRT
51-2-4-F5-CL11_2	182	PRT
51-27-4-F2-CLO_2	183	PRT
51-34-3-F8-CLO_2	184	PRT

57-1-4-E2-CL1_2	185	PRT
57-19-2-G8-CL2_1	186	PRT
57-27-3-G10-CL2_2	187	PRT
58-33-3-B4-CL1_2	188	PRT
58-34-3-C9-CL1_2	189	PRT
58-4-4-G2-CL2_1	190	PRT
58-48-1-G3-CL2_4	191	PRT
58-6-1-H4-CL1_1	192	PRT
60-12-1-E11-CL1_2	193	PRT
65-4-4-H3-CL1_1	194	PRT
74-5-1-E4-CL1_2	195	PRT
76-13-3-A9-CL1_2	196	PRT
76-16-1-D6-CL1_1	197	PRT
76-28-3-A12-CL1_5	198	PRT
76-42-2-F3-CLO_1	199	PRT
77-16-4-G3-CL1_3	200	PRT
77-39-4-H4-CL11_4	201	PRT
78-24-3-H4-CL2_1	202	PRT
78-27-3-D1-CL1_6	203	PRT
78-28-3-D2-CL0_2	204	PRT
78-7-1-G5-CL2_6	205	PRT
84-3-1-G10-CL11_6	206	PRT
58-48-4-E2-CLO_1	207	PRT
23-12-2-G6-CL1_2	208	PRT
25-8-4-B12-CL0_5	209	PRT
26-44-3-C5-CL2_1	210	PRT
27-1-2-B3-CL0_3	211	PRT
30-12-3-G5-CL0_1	212	PRT
33-106-2-F10-CL1_3	213	PRT
33-28-4-D1-CL0_1	214	PRT
33-31-3-C8-CL2_1	215	PRT
48-24-1-D2-CL3_2	216	PRT
48-46-4-A11-CL1_4	217	PRT
51-1-4-C1-CL0_2	218	PRT
51-39-3-H2-CL1_2	219	PRT
51-42-3-F9-CL1_1	220	PRT
51-5-3-G2-CL0_4	221	PRT

57-18-4-H5-CL2_1	222	PRT
76-23-3-G8-CL1_1	223	PRT
76-23-3-G8-CL1_3	224	PRT
78-8-3-E6-CLO_1	225	PRT
19-10-1-C2-CL1_3	226	PRT
33-11-1-B11-CL1_2	227	PRT
33-113-2-B8-CL1_2	228	PRT
33-19-1-C11-CL1_1	229	PRT
33-61-2-F6-CLO_2	230	PRT
47-4-4-C6-CL2_2	231	PRT
48-54-1-G9-CL1_1	232	PRT
51-43-3-G3-CLO_1	233	PRT
55-1-3-D11-CLO_1	234	PRT
58-14-2-D3-CL1_2	235	PRT
58-35-2-B6-CL2_3	236	PRT
76-18-1-F6-CL1_1	237	PRT
76-23-3-G8-CL2_2	238	PRT
76-30-3-B7-CL1_1	239	PRT
78-21-3-G7-CL2_1	240	PRT
58-45-4-B11-CL13_2	241	PRT
20-6-1-D11-FL2	242	DNA
20-8-4-A11-FL2	243	DNA
22-6-2-C1-FL2	244	DNA
22-11-2-H9-FL1	245	DNA
23-8-3-B1-FL1	246	DNA
24-3-3-C6-FL1	247	DNA
24-4-1-H3-FL1	248	DNA
26-45-2-C4-FL2	249	DNA
26-48-1-H10-FL1	250	DNA
26-49-1-A5-FL2	251	DNA
30-6-4-E3-FL3	252	DNA
33-6-1-G11-FL1	253	DNA
33-8-1-A3-FL2	254	DNA
33-11-3-C6-FL1	255	DNA
33-14-4-E1-FL1	256	DNA
33-21-2-D5-FL1	257	DNA
33-26-4-E10-FL1	258	DNA

00.07 + 514 514	250	DNA
33-27-1-E11-FL1	259	DNA
33-28-4-D1-FL1	260	DNA
33-28-4-E2-FL2	261	DNA
33-30-4-C4-FL1	262	DNA
33-35-4-F4-FL1	263	DNA
33-36-3-F2-FL2	264	DNA
33-52-4-F9-FL2	265	DNA
33-52-4-H3-FL1	266	DNA
33-59-1-B7-FL1	267	DNA
33-71-1-A8-FL1	268	DNA
33-72-2-B2-FL1	269	DNA
33-105-2-C3-FL1	270	DNA
33-107-4-C3-FL1	271	DNA
33-110-2-G4-FL1	272	DNA
47-7-4-D2-FL2	273	DNA
47-10-2-G12-FL1	274	DNA
47-14-3-D8-FL1	275	DNA
47-18-3-C2-FL1	276	DNA
47-18-3-G5-FL2	277	DNA
47-18-4-E3-FL2	278	DNA
48-3-1-H9-FL3	279	DNA
48-4-2-H3-FL1	280	DNA
48-6-1-C9-FL1	281	DNA
48-7-4-H2-FL2	282	DNA
48-8-1-D8-FL3	283	DNA
48-13-3-H8-FL1	284	DNA
48-19-3-A7-FL1	285	DNA
48-19-3-G1-FL1	286	DNA
48-25-4-D8-FL1	287	DNA
48-21-4-H4-FL1	288	DNA
48-26-3-B8-FL2	289	DNA
48-29-1-E2-FL1	290	DNA
48-31-3-F7-FL1	291	DNA
48-47-3-A5-FL1	292	DNA
51-1-1-G12-FL1	293	DNA
51-1-4-E9-FL3	294	DNA
51-1-4-E9-FL2	295	DNA
	<u> </u>	

296	DNA
297	DNA
298	DNA
299	DNA
300	DNA
301	DNA
302	DNA
303	DNA
304	DNA
305	DNA
306	DNA
307	DNA
308	DNA
309	DNA
310	DNA
311	DNA
312	DNA
313	DNA
314	DNA
315	DNA
316	DNA
317	DNA
318	DNA
319	DNA
320	DNA
321	DNA
322	DNA
323	DNA
324	DNA
325	DNA
326	DNA
327	DNA
328	DNA
329	DNA
330	DNA
331	DNA
332	DNA
	298 299 300 301 302 303 304 305 306 307 308 309 310 311 312 313 314 315 316 317 318 319 320 321 322 323 324 325 326 327 328 329 330 331

·		•
65-4-4-H3-FL1	333	DNA
74-3-1-89-FL1	334	DNA
76-4-1-G5-FL1	335	DNA
76-7-3-A12-FL1	336	DNA
76-16-4-C9-FL3	337	DNA
76-30-3-B7-FL1	338	DNA
77-5-1-C2-FL1	339	DNA
77-5-4-E7-FL1	340	DNA
77-11-1-A3-FL1	341	DNA
77-16-3-D7-FL1	342	DNA
77-16-4-G3-FL1	343	DNA
77-25-1-A6-FL1	344	DNA
77-26-2-F2-FL3	345	DNA
78-6-2-E3-FL2	346	DNA
78-7-1-G5-FL2	347	DNA
78-16-2-C2-FL1	348	DNA
78-18-3-B4-FL3	349	DNA
78-20-1-G11-FL1	350	DNA
78-22-3-E10-FL1	351	DNA
78-24-2-B8-FL1	352	DNA
78-24-3-A8-FL1	353	DNA
78-24-3-H4-FL2	354	DNA
78-25-1-F11-FL1	355	DNA
78-26-1-B5-FL1	356	DNA
78-27-3-D1-FL1	357	DNA
78-29-1-B2-FL1	358	DNA
78-29-4-B6-FL1	359	DNA
14-1-3-E6-FL1	360	DNA
30-9-1-G8-FL2	361	DNA
33-10-4-H2-FL2	362	DNA -
33-10-4-H2-FL1	363	DNA
74-10-3-C9-FL2	364	DNA
33-97-4-G8-FL3	365	DNA
33-97-4-G8-FL2	366	DNA
33-104-4-H4-FL1	367	DNA
47-2-3-B3-FL1	368	DNA
47-37-4-G11-FL1	369	DNA

57-25-1-F10-FL2	370	DNA
58-19-3-D3-FL1	371	DNA
58-34-3-C9-FL2	372	DNA
58-48-4-E2-FL2	373	DNA
76-21-1-C4-FL1	374	DNA
78-26-2-H7-FL1	375	DNA
77-20-2-E11-FL1	376	DNA
47-1-3-F7-FL2	377	DNA
20-6-1-D11-FL2	378	PRT
20-8-4-A11-FL2	379	PRT
22-6-2-C1-FL2	380	PRT
22-11-2-H9-FL1	381	PRT
23-8-3-B1-FL1	382	PRT
24-3-3-C6-FL1	383	PRT
24-4-1-H3-FL1	384	PRT
26-45-2-C4-FL2	385	PRT
26-48-1-H10-FL1	386	PRT
26-49-1-A5-FL2	387	PRT
30-6-4-E3-FL3	388	PRT
33-6-1-G11-FL1	389	PRT
33-8-1-A3-FL2	390	PRT
33-11-3-C6-FL1	391	PRT
33-14-4-E1-FL1	392	PRT
33-21-2-D5-FL1	393	PRT
33-26-4-E10-FL1	394	PRT
33-27-1-E11-FL1	395	PRT
33-28-4-D1-FL1	396	PRT
33-28-4-E2-FL2	397	PRT
33-30-4-C4-FL1	398	PRT
33-35-4-F4-FL1	399	PRT
33-36-3-F2-FL2	400	PRT
33-52-4-F9-FL2	401	PRT
33-52-4-H3-FL1	402	PRT
33-59-1-B7-FL1	403	PRT
33-71-1-A8-FL1	404	PRT
33-72-2-B2-FL1	405	PRT
33-105-2-C3-FL1	406	PRT

		50 -
33-107-4-C3-FL1	407	PRT
33-110-2-G4-FL1	408	PRT
47-7-4-D2-FL2	409	PRT
47-10-2-G12-FL1	410	PRT
47-14-3-D8-FL1	411	PRT
47-18-3-C2-FL1	412	PRT
47-18-3-G5-FL2	413	PRT
47-18-4-E3-FL2	414	PRT
48-3-1-H9-FL3	415	PRT
48-4-2-H3-FL1	416	PRT
48-6-1-C9-FL1	417	PRT
48-7-4-H2-FL2	418	PRT
48-8-1-D8-FL3	419	PRT
48-13-3-H8-FL1	420	PRT
48-19-3-A7-FL1	421	PRT
48-19-3-G1-FL1	422	PRT
48-25-4-D8-FL1	423	PRT
48-21-4-H4-FL1	424	PRT
48-26-3-B8-FL2	425	PRT
48-29-1-E2-FL1	426	PRT
48-31-3-F7-FL1	427	PRT
48-47-3-A5-FL1	428	PRT
51-1-1-G12-FL1	429	PRT
51-1-4-E9-FL3	430	PRT
51-1-4-E9-FL2	431	PRT
51-2-1-E10-FL1	432	PRT
51-2-3-F10-FL1	433	PRT
51-2-4-F5-FL1	434	PRT
51-3-3-B10-FL2	435	PRT
51-3-3-B10-FL3	436	PRT
51-7-3-G3-FL1	437	PRT
51-10-3-D11-FL1	438	PRT
51-11-3-D5-FL1	439	PRT
51-13-1-F7-FL3	440	PRT
51-15-4-H10-FL1	441	PRT
51-17-4-A4-FL1	442	PRT
51-18-1-C3-FL1	443	PRT

		·
51-25-3-F3-FL1	444	PRT
51-27-1-E8-FL1	445	PRT
51-28-2-G1-FL2	446	PRT
51-39-3-H2-FL1	447	PRT
51-42-3-F9-FL1	448	PRT
51-44-4-H4-FL1	449	PRT
55-1-3-H10-FL1	450	PRT
55-5-4-A6-FL1	451	PRT
58-26-3-D1-FL1	452	PRT
57-18-1-D5-FL1	453	PRT
57-27-3-A11-FL1	454	PRT
57-27-3-G10-FL2	455	PRT
58-10-3-D12-FL1	456	PRT
58-11-1-G10-FL1	457	PRT
58-11-2-G8-FL2	458	PRT
58-36-3-A9-FL2	459	PRT
58-38-1-A2-FL2	460	PRT
58-38-1-E5-FL1	461	PRT
58-44-2-B3-FL3	462	PRT
58-45-3-H11-FL1	463	PRT
58-53-2-B12-FL2	464	PRT
59-9-4-A10-FL1	465	PRT
60-16-3-A6-FL1	466	PRT
60-17-3-G8-FL2	467	PRT
62-5-4-B10-FL1	468	PRT
65-4-4-H3-FL1	469	PRT
74-3-1-B9-FL1	470	PRT
76-4-1-G5-FL1	471	PRT
76-7-3-A12-FL1	472	PRT
76-16-4-C9-FL3	473	PRT
76-30-3-B7-FL1	474	PRT
77-5-1-C2-FL1	475	PRT
77-5-4-E7-FL1	476	PRT
77-11-1-A3-FL1	477	PRT
77-16-3-D7-FL1	478	PRT
77-16-4-G3-FL1	479	PRT
77-25-1-A6-FL1	480	PRT

77-26-2-F2-FL3	481	PRT	
78-6-2-E3-FL2	482	PRT	
78-7-1-G5-FL2	483	PRT	
78-16-2-C2-FL1	484	PRT	
78-18-3-B4-FL3	485	PRT	
78-20-1-G11-FL1	486	PRT	
78-22-3-E10-FL1	487	PRT	
78-24-2-B8-FL1	488	PRT	
78-24-3-A8-FL1	489	PRT	
78-24-3-H4-FL2	490	PRT	
78-25-1-F11-FL1	491	PRT	
78-26-1-B5-FL1	492	PRT	
78-27-3-D1-FL1	493	PRT	
78-29-1-B2-FL1	494	PRT	
78-29-4-B6-FL1	495	PRT	
14-1-3-E6-FL1	496	PRT	
30-9-1-G8-FL2	497	PRT	
33-10-4-H2-FL2	498	PRT	
33-10-4-H2-FL1	499	PRT	
74-10-3-C9-FL2	500	PRT	
33-97-4-G8-FL3	501	PRT	
33-97-4-G8-FL2	502	PRT	
33-104-4-H4-FL1	503	PRT	
47-2-3-B3-FL1	504	PRT	
47-37-4-G11-FL1	505	PRT	
57-25-1-F10-FL2	506	PRT	
58-19-3-D3-FL1	507	PRT	
58-34-3-C9-FL2	508	PRT	
58-48-4-E2-FL2	509	PRT	
76-21-1-C4-FL1	510	PRT	
78-26-2-H7-FL1	511	PRT	
77-20-2-E11-FL1	512	PRT	
47-1-3-F7-FL2	513	PRT	

-136-

TABLE VIII

ID	Locations	PROSITE Signature Name		
195	110-121	Aldehyde dehydrogenases csyteine active site		
221	28-37	ATP synthase alpha and beta subunits signature		
223	171-181	Regulator of chromosome condensation (RCC1) signature 2		
225	- 90-112	Phosphatidylethanolamine-binding protein family signature		
226	10-34	Protein kinases ATP-binding region signature		

WHAT IS CLAIMED IS:

10

- 1. A purified or isolated nucleic acid comprising the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary thereto.
- 2. A purified or isolated nucleic acid comprising at least 10 consecutive bases of the sequence of one of SEQ ID NOs: 40-140 and 242-377 or one of the sequences complementary thereto.
 - 3. A purified or isolated nucleic acid comprising the full coding sequences of one of SEQ ID NOs: 40, 42-44, 46, 48, 49, 51, 53, 60, 62-72, 76-78, 80-83, 85-88, 90, 93, 94, 97, 99-102, 104, 107-125, 127, 132, 135-138, 140 and 242-377wherein the full coding sequence comprises the sequence encoding signal peptide and the sequence encoding mature protein.
 - 4. A purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode a mature protein.
- 5. A purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40, 42-46, 48, 49, 51, 53, 57, 60, 62-73, 76-78, 80-83, 85-88, 90, 93-95, 97, 99-102, 104, 107-125, 127, 128, 130, 132, 134-140 and 242-377 which encode the signal peptide.
 - 6. A purified or isolated nucleic acid encoding a polypeptide having the sequence of one of the sequences of SEQ ID NOs: 141-241 and 378-513.
- 7. A purified or isolated nucleic acid encoding a polypeptide having the sequence of a mature protein included in one of the sequences of SEO ID NOs: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-20 189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.
 - 8. A purified or isolated nucleic acid encoding a polypeptide having the sequence of a signal peptide included in one of the sequences of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-226, 228, 229, 231, 233, 235-241, and 378-513.
 - 9. A purified or isolated protein comprising the sequence of one of SEQ ID NOs: 141-241 and 378-513.
- 25 10. A purified or isolated polypeptide comprising at least 10 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513.
 - 11. An isolated or purified polypeptide comprising a signal peptide of one of the polypeptides of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-226, 228, 229, 231, 233, 235-241, and 378-513.
- 30 12. An isolated or purified polypeptide comprising a mature protein of one of the polypeptides of SEQ ID NOs: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.
 - 13. A method of making a protein comprising one of the sequences of SEQ ID NO: 141-241 and 378-513, comprising the steps of:

cDNA.

obtaining a cDNA comprising one of the sequences of sequence of SEQ ID NO: 40-140 and 242-377; inserting said cDNA in an expression vector such that said cDNA is operably linked to a promoter; and introducing said expression vector into a host cell whereby said host cell produces the protein encoded by said

- 5 14. The method of Claim 13, further comprising the step of isolating said protein.
 - 15. A protein obtainable by the method of Claim 14.
 - 16. A host cell containing a recombinant nucleic acid of Claim 1.
 - 17. A purified or isolated antibody capable of specifically binding to a protein having the sequence of one of SEQ ID NOs: 141-241 and 378-513.
- 10 18. In an array of polynucleotides of at least 15 nucleotides in length, the improvement comprising inclusion in said array of at least one of the sequences of SEQ ID NOs: 40-140 and 242-377, or one of the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or a fragment thereof of at least 15 consecutive nucleotides.
- 19. A purified or isolated nucleic acid of at least 15 bases capable of hybridizing under stringent

 15 conditions to the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary to one of the sequences of SEQ ID NOs: 40-140 and 242-377.
 - 20. A purified or isolated antibody capable of binding to a polypeptide comprising at least 10 consecutive amino acids of the sequence of one of SEQ ID NOs: 141-241 and 378-513.

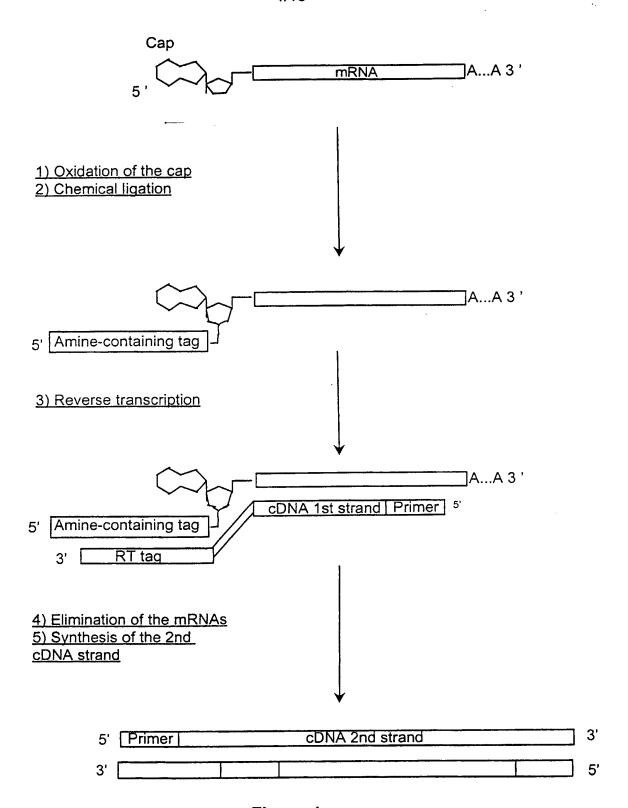


Figure 1

Minimum signal peptide score	false positive rate	false negative rate	proba(0.1)	proba(0.2)
3,5	0,121	0,036	0,467	0,664
4	0,096	0,06	0,519	0,708
4,5	0,078	0,079	0,565	0,745
5	0,062	0,098	0,615	0,782
5,5	0,05	0,127	0,659	0,813
6	0,04	0,163	0,694	0,836
6,5	0,033	0,202	0,725	0,855
7	0,025	0,248	0,763	0,878
7,5	0,021	0,304	0,78	0,889
8	0,015	0,368	0,816	0,909
8,5	0,012	0,418	0,836	0,92
-9	• 0,009	0,512	0,856	0,93
9,5	0,007	0,581	0,863	0,934
10	0,006	0,679	0,835	0,919

influence of minimum score on signal peptide recognition

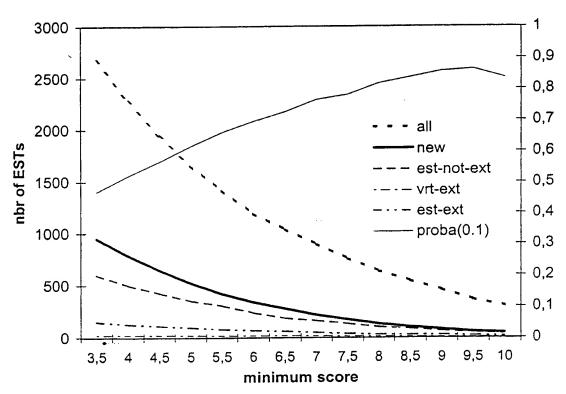
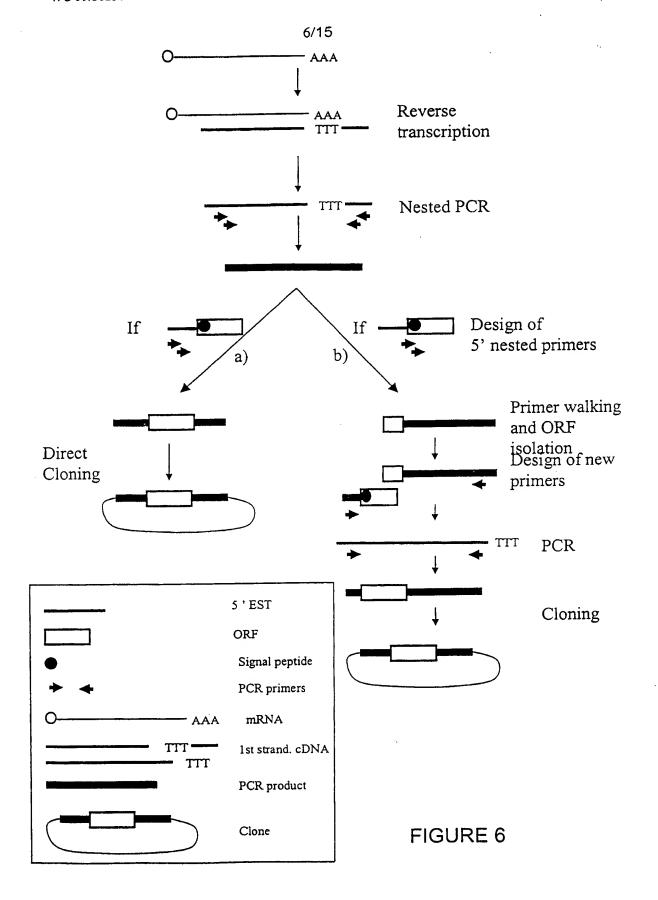
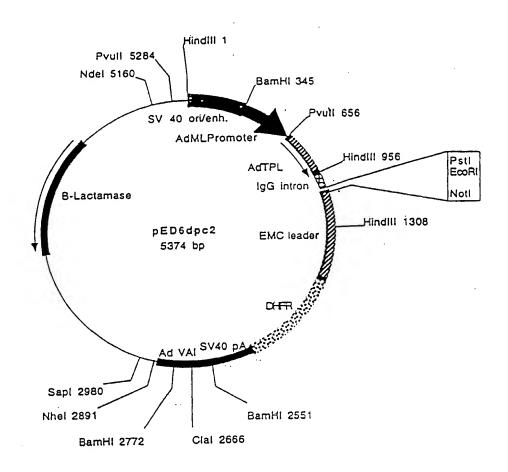


FIGURE 3

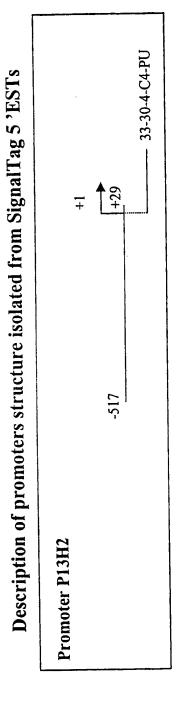
Minimum signal peptide score	All ESTs	New ESTs	ESTs matching public EST closer than 40 bp from beginning	ESTs extending known mRNA more than 40 bp	ESTs extending public EST more than 40 bp
3,5	2674	947	599	23	150
4	2278	784	499	23	126
4,5	1943	647	425	22	112
5	1657	523	353	21	96
5,5	1417	419	307	19	80
6	1190	340	238	18	68
6,5	1035	280	186	18	60
7	893	219	161	15	48
7,5	753	173	132	12	36
8	636	133	101	11	29
8,5	543	104	83	8	26
9	456	81	63	6	24
9,5	364	57	48	6	18
10	303	47	35	6	15

Tissue	All ESTs	New ESTs	ESTs matching public EST closer than 40 bp from beginning	ESTs extending known mRNA more than 40 bp	ESTs extending public EST more than 40 bp
Brain	329	131	75	3	24
Cancerous prostate	134	40	37	1	6
Cerebellum	17	9	1	0	6
Colon	21	11	4	0	0
Dystrophic muscle	41	18	8	0	1
Fetal brain	70	37	16	0	1
Fetal kidney	227	116	46	1	19
Fetal liver	13	7	2	0	0
Heart	30	15	7	0	1
Hypertrophic prostate	86	23	22	2	2
Kidney	10	7	3	0	0
Large intestine	21	8	4	0	1
Liver	23	9	6	0	0
Lung	24	12	4	0	1
Lung (cells)	57	38	6	0	4
Lymph ganglia	163	60	23	2	12
Lymphocytes	23	6	4	0	2
Muscle	33	16	6	0	4
Normal prostate	181	61	45	7	11
Ovary	90	57	12	1	2
Pancreas	48	11	6	0	1
Placenta	24	5	1	0	0
Prostate	34	16	4	0	2
Spleen	56	28	10	0	1
Substantia nigra	108	47	27	. 1	6
Surrenals	15	3	3	1	0
Testis	131	68	25	1	8
Thyroid	17	8	2	0	2 3
Umbilical cord	55	17	12	1	
Uterus	28	15	3	0	2
Non tissue-specific	568	48	177	2	28
Total	2677	947	601	23	150





Plasmid name: pED6dpc2 Plasmid size: 5374 bp 8/15



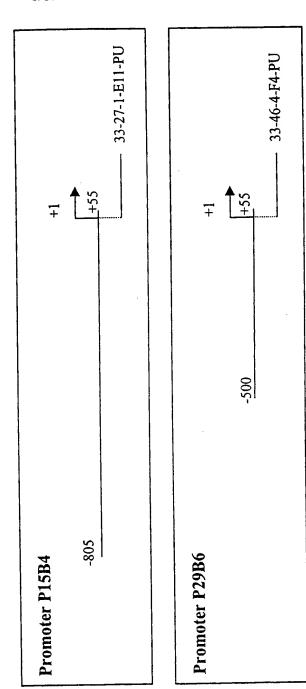


FIGURE 8

9/15

Description of Transcription Factor Binding Sites present on promoters isolated from SignalTag sequences

Promoter sequence P13H2 (546 bp):

Matrix	Position	Orientation	Score	Length	Sequence
CMYB_01	-502	+	0.983	9	TGTCAGTTG
MYOD_Q6	-501	- .	0.961	10	CCCAACTGAC
S8_01	-444	•	0.960	11	AATAGAATTAG
S8 01	-425	+	0.966	11	AACTAAATTAG
DELTAEF1_01	-390	•	0.960	11	GCACACCTCAG
GATA_C	-364	•	0.964	11	AGATAAATCCA
CMYB_01	-349	+	0.958	9	CTTCAGTTG
GATA1_02	-343	+	0.959	14	TTGTAGATAGGACA
GATA_C	-339	+	0.953	11	AGATAGGACAT
TAL1ALPHAE47 01	-235	+	0.973	16	CATAACAGATGGTAAG
TAL1BETAE47_01	-235	+	0.983	16	CATAACAGATGGTAAG
TAL1BETAITF2_01	-235	+	0.978	16	CATAACAGATGGTAAG
MYOD_Q6	-232	•	0.954	10	ACCATCTGTT
GATA1_04	-217	-	0.953	13	TCAAGATAAAGTA
IK1_01	-126	+	0.963	13	AGTTGGGAATTCC
IK2_01	-126	+	0.985	12	AGTTGGGAATTC
CREL_01	-123	+	0.962	10	TGGGAATTCC
GATA1_02	-96	+	0.950	14	TCAGTGATATGGCA
SRY_02	-41	-	0.951	12	TAAAACAAAACA
E2F_02	-33	+	0.957	8	TTTAGCGC
MZF1_01	-5	•	0.975	8	TGAGGGGA

Promoter sequence P15B4 (861bp):

Matrix	Position	Orientation	Score	Length	Sequence
NFY_Q6	-748	-	0.956	11	GGACCAATCAT
MZF1_01	-738	+	0.962	8	CCTGGGGA
CMYB_01	-684	, +	0.994	. 9	TGACCGTTG
VMYB_02	-682	•	0.985	9	TCCAACGGT
STAT_01	-673	+	0.968	9	TTCCTGGAA
STAT_01	-673	•	0.951	9	TTCCAGGAA
MZF1_01	-556	-	0.956	8	TTGGGGGA
IK2_01	-451	+	0.965	12	GAATGGGATTTC
MZF1_01	-424	+	0.986	8	AGAGGGGA
SRY_02	-398	-	0.955	12	GAAAACAAAACA
MZF1_01	-216	+	0.960	8	GAAGGGGA
MYOD_Q6	-190	+	0.981	10	AGCATCTGCC
DELTAEF1_01	-176	+	0.958	11	TCCCACCTTCC
S8_01	5	-	0.992	11	GAGGCAATTAT
MZF1_01	16	-	0.986	8	AGAGGGGA

Promoter sequence P29B6 (555 bp):

Matrix	Position (Orientation	Score	Length	Sequence
ARNT_01	-311	+	0.964	16	GGACTCACGTGCTGCT
NMYC_01	-309	+	0.965	12	ACTCACGTGCTG
USF_01	-309	+	0.985	12	ACTCACGTGCTG
USF_01	-309	-	0.985	12	CAGCACGTGAGT
NMYC_01	-309	-	0.956	12	CAGCACGTGAGT
MYCMAX_02	-309	•	0.972	12	CAGCACGTGAGT
USF_C	-307	+	0.997	8	TCACGTGC
USF_C	-307	•	0.991	8	GCACGTGA :
MZF1_01	-292	•	0.968	8	CATGGGGA
ELK1_02	-105	+	0.963	14	CTCTCCGGAAGCCT
CETS1P54_01	-102	+	0.974	10	TCCGGAAGCC
AP1_Q4	-42	•	0.963	11	AGTGACTGAAC
AP1FJ_Q2	-42	•	0.961	11	AGTGACTGAAC
PADS_C	45	+	1.000	9	TGTGGTCTC

Figure 9

10/15

100.0% identity in 125 aa overlap 40 50 60 20 30 SEQ ID NO: 217 MADEELEALRRQRLAELQAKHGDPGDAAQQEAKHREAEMRNSILAQVLDQSARARLSNLA SEQ ID NO: 516 MADEELEALRRQRLAELQAKHGDPGDAAQQEAKHREAEMRNSILAQVLDQSARARLSNLA 40 20 30 120 70 80 90 100 110 SEQ ID NO: 217 LVKPEKTKAVENYLIQMARYGQLSEKVSEQGLIEILKKVSQQTEKTTTVKFNRRKVMDSD SEQ ID NO: 516 LVKPEKTKAVENYLIQMARYGQLSEKVSEQGLIEILKKVSQQTEKTTTVKFNRRKVMDSD 80 90 100 110 70 SEQ ID NO: 217 EDDDY ::::X SEQ ID NO: 516 EDDDY

11/15

CLUSTAL W(1.5) multiple sequence alignment

SEQ ID NO: 517 SEQ ID NO: 232 SEQ ID NO: 174 SEQ ID NO: 175	MFCPLKLILLPVLLDYSLGLNDLNVSPPELTVHVGDSALMGCVFQSTEDKCIFKIDWTLS
SEQ ID NO: 517 SEQ ID NO: 232 SEQ ID NO: 174 SEQ ID NO: 175	PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDNLCNDGSLLLQDVQDVE PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDILCNDGSLLLQDVQEADQGTYICEIRL PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDNLCNDGSLLLQDVQEADQGTYICEIRL PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDILCNDGSLLLQDVQEADQGTYICEIRL
SEQ ID NO: 517 SEQ ID NO: 232 SEQ ID NO: 174 SEQ ID NO: 175	KGESQVFKKAVVLHVLPEEPKGTQMLTKGESQVFKKAVVLHVLPEEPKELMVHVGGLIQMGCVFQSTEVKHVTKVEWIFSGRRAKEEKGESQVFKKAVVLHVLPEEPKELMVHVGGLIQMGCVFQSTEVKHVTKVEWIFSGRRAK
SEQ ID NO: 517 SEQ ID NO: 232 SEQ ID NO: 174 SEQ ID NO: 175	IVFRYYHKLRMSAEYSQSWGHFQNRVNLVGDIFRNDGSIMLQGVRESDGGNYTCSIHLGN VTRRKHHCVREGSG
SEQ ID NO: 517 SEQ ID NO: 232 SEQ ID NO: 174 SEQ ID NO: 175	LVFKKTIVLHVSPEEPRTLVTPAALRPLVLGGNQLVIIVGIVCATILLLPVLILIVKKTC
SEQ ID NO: 517 SEQ ID NO: 232 SEQ ID NO: 174 SEQ ID NO: 175	GNKSSVNSTVLVKNTKKTNP

12/15

99.6% identity in 225 aa overlap SEQ ID NO: 515 PTAVQKEEARQDVEALLSRTVRTQILTGKELRVATQEKEGSSGRCMLTLLGLSFILAGLI LRVATQEKEGSSGRCMLTLLGLSFILAGLI SEQ ID NO: 231 SEQ ID NO: 515 VGGACIYKYFMPKSTIYRGEMCFFDSEDPANSLRGGEPNFLPVTEEADIREDDNIAIIDV SEQ ID NO: 231 VGGACIYKYFMPKSTIYRGEMCFFDSEDPANSLRGGEPNFLPVTEEADIREDDNIAIIDV SEQ ID NO: 515 PVPSFSDSDPAAIIHDFEKGMTAYLDLLLGNCYLMPLNTSIVMPPKNLVELFGKLASGRY SEQ ID NO: 231 PVPSFSDSDPAAIIHDFEKGMTAYLDLLLGICYLMPLNTSIVMPPKNLVELFGKLASGRY SEQ ID NO: 515 LPQTYVVREDLVAVEEIRDVSNLGIFIYQLCNNRKSFRLRRRDLLLGFNKRAIDKCWKIR SEQ ID NO: 231 LPQTYVVREDLVAVEEIRDVSNLGIFIYQLCNNRKSFRLRRRDLLLGFNKRAIDKCWKIR 170 180 SEQ ID NO: 515 HFPNEFIVETKICQE :::::::::::::::: SEQ ID NO: 231 HFPNEFIVETKICQE

13/15

99.7% identity in 353 aa overlap MERGLKSADPRDGTGYTGWAGIAVLYLHLY SEQ ID NO:196 SEQ ID NO:518 LAEGYFDAAGRLTPEFSQRLTNKIRELLQQMERGLKSADPRDGTGYTGWAGIAVLYLHLY SEQ ID NO:196 DVFGDPAYLQLAHGYVKQSLNCLTKRSITFLCGDAGPLAVAAVLYHKMNNEKQAEDCITR SEQ ID NO:518 DVFGDPAYLQLAHGYVKQSLNCLTKRSITFLCGDAGPLAVAAVLYHKMNNEKQAEDCITR SEQ ID NO:196 LIHLNKIDPHAPNEMLYGRIGYIYALLFVNKNFGVEKTPQSHIQQICETILTSGENLARK SEQ ID NO:518 LIHLNKIDPHAPNEMLYGRIGYIYALLFVNKNFGVEKIPQSHIQQICETILTSGENLARK SEQ ID NO:196 RNFTAKSPLMYEWYQEYYVGAAHGLAGIYYYLMQPSLQVSQGKLHSLVKPSVDYVCQLKF SEQ ID NO:518 RNFTAKSPLMYEWYQEYYVGAAHGLAGIYYYLMQPSLQVSQGKLHSLVKPSVDYVCQLKF SEQ ID NO:196 PSGNYPPCIGDNRDLLVHWCHGAPGVIYMLIQAYKVFREEKYLCDAYQCADVIWQYGLLK SEQ ID NO:518 PSGNYPPCIGDNRDLLVHWCHGAPGVIYMLIQAYKVFREEKYLCDAYQCADVIWQYGLLK SEQ ID NO:196 KGYGLCHGSAGNAYAFLTLYNLTQDMKYLYRACKFAEWCLEYGEHGCRTPDTPFSLFEGM SEQ ID NO:518 KGYGLCHGSAGNAYAFLTLYNLTQDMKYLYRACKFAEWCLEYGEHGCRTPDTPFSLFEGM SEO ID NO:196 AGTIYFLADLLVPTKARFPAFEL SEQ ID NO:518 AGTIYFLADLLVPTKARFPAFEL

14/15

98.5% identity in 194 aa overlap 120 130 100 110 90 SEQ ID NO:519 ARNLPPLTDAQKNKLRHLSVVTLAAKVKCIPYAVLLEALALRNVRQLEDLVIEAVYADVL SEQ ID NO:158 ARNLPPLTEAQKNKLRHLSVVTLAAKVKCIPYAVLLEALALRNVRQLEDLVIEAVYADVL 80 90 100 70 180 160 170 190 150 SEQ ID NO:519 RGSLDQRNQRLEVDYSIGRDIQRQDLSAIAQTLQEWCVGCEVVLSGIEEQVSRANQHKEQ SEQ ID NO:158 RGSLDQRNQRLEVDYSIGRDIQRQDLSAIARTLQEWCVGCEVVLSGIEEQVSRANQHKEQ 130 140 210 230 240 250 220 SEQ ID NO:519 QLGLKQQIESEVANLKKTIKVTTAAAAAATSQDPEQHLTELREPASGTNQRQPSKKASKG SEQ ID NO:158 QLGLKQQIESEVANLKKTIKVTTAAAAAATSQDPEQHLTELREPAPGTNQRQPSKKASKG 200 210 190 270 SEQ ID NO:519 KGLRGSAKIWSKSN SEQ ID NO:158 KGLRGSAKIWSKSN 88.7% identity in 62 aa overlap 40 10 20 30 SEQ ID NO:519 MSAEVKVTGQNQEQFLLLAKSAKGAALATLIHQVLEAPGVYVFGELLDMPNVRELAESDF SEQ ID NO:158 MSAEVKVTGQNQEQFLLLAKSAKGAALATLIHQVLEAPGVYVFGELLDMPNVRELXARNL 20 30 10

SEQ ID NO:519 AS

SEQ ID NO:158 PP

15/15

68.9% identity	y in 74 aa over]	.ap				•.
	10 MIARRNPVPLRFL	20	30	40	50	TAGLHR
SEQ ID NO:226			:::::::	::::::::	:: :::::	:::::
SEQ ID NO:514	MMTGRQGRATFQFLE	PDEARSLPPP 20	KLTDPRLAFVO 30	3FLGYCSGL11 40	50	60
	60 70					
	QLLYITAFFLLDIII					
SEQ ID NO:514	QLLYITSFVFVGYYI		VRDHDMFSYII 90	KSHPEDFPEK 100	DKKTYGEVFE 110	EFHPVR 120
	70	00	50	200		- - -

<120> Extended cDNAS for Secreted Proteins

<130> GENSET.019A

<160> 519

<170> Patent.pm

<210> 1

<211> 47

<212> RNA

<213> Artificial Sequence

<220>

<221> In vitro transcription product

<221> modified_base

<222> (1)...(1)

<223> m7g

<400> 1

ngcauccuac ucccauccaa uuccacccua acuccuccca ucuccac

47

<210> 2

<211> 46

<212> RNA (

<213> Artificial Sequence

<220>

<223> In vitro transcription product

<400> 2

gcauccuacu cccauccaau uccacccuaa cuccucccau cuccac

46

<210> 3

<211> 25

<212> DNA

<213> Artificial Sequence

<220>

<223> In vitro transcription product

<400> 3

atcaagaatt cgcacgagac catta

25

<210> 4

<211> 25

<212> DNA

<213> Artificial Sequence

<220>		
<223>	Oligonucleotide	
400-		
<400>	tctc gtgcgaattc ttgat	25
Laacyy		
<210>		
<211><212>		
	Artificial Sequence	
12207	•	
<220>		
<223>	Oligonucleotide	
<400>	E C	
	agac caacgtcaag gccgc	25
CCGGCG		
<210> <211>		
<211>		
	Artificial Sequence	
<220>		
<223>	Oligonucleotide	
<400>	6	
	agcag gcagtggctt aggag	25
<210>	7	
<211>		
<212>		
<213>	Artificial Sequence	
.000-		
<220>	Oligonucleotide	
~~~~		
<400>	7	25
agtga	ttoot gotaotttgg atggo	25
<210>	8	
<211>		
<212>		
<213>	Artificial Sequence	
<220>		
	Oligonucleotide	
<400>		25
gcttg	gtctt gttctggagt ttaga	

<211> 25	
<212> DNA <213> Artificial Sequence	
<220> <223> Oligonucleotide	
<400> 9	
tccagaatgg gagacaagcc aattt	25
<210> 10 <211> 25	
<212> DNA	
<213> Artificial Sequence	
<220>	
<223> Oligonucleotide	
<400> 10	25
agggaggagg aaacagcgtg agtcc	
	•
<210> 11 <211> 25	
<211> 25 <212> DNA	
<213> Artificial Sequence	
<220>	
<223> Oligonucleotide	
<400> 11	25
atgggaaagg aaaagactca tatca	
<210> 12	
<211> 25 <212> DNA	
<213> Artificial Sequence	
<220>	
<223> Oligonucleotide	
<400> 12	25
agcagcaaca atcaggacag cacag	
-	
<210> 13	
<211> 25 <212> DNA	
<212> DNA <213> Artificial Sequence	
<220>	
<223> Oligonucleotide	
<400> 13	25
atcaagaatt cgcacgagac catta	

WO 99/31236

```
<210> 14
<211> 67
<212> DNA
<213> Artificial Sequence
<220>
<223> Oligonucleotide
<400> 14
atcgttgaga ctcgtaccag cagagtcacg agagagacta cacggtactg gtttttttt
                                                                        60
                                                                         67
tttttvn
<210> 15
<211> 29
<212> DNA
<213> Artificial Sequence
<220>
<223> Oligonucleotide
<400> 15
                                                                         29
ccagcagagt cacgagagag actacacgg
<210> 16
<211> 25
<212> DNA
<213> Artificial Sequence
<220>
<223> Oligonucleotide
<400> 16
                                                                         25
cacgagagag actacacggt actgg
<210> 17
<211> 526
<212> DNA
<213> Homo sapiens
<220>
<221> misc_feature
<222> complement (261..376)
<223> blastn
<221> misc_feature
<222> complement (380..486)
 <223> blastn
 <221> misc_feature
 <222> complement(110..145)
 <223> blastn
 <221> misc_feature
 <222> complement (196..229)
 <223> blastn
```

-5-

60

209

257

305

354

414

474

526

<221> sig peptide <222> 90..140 <223> Von Heijne matrix <400> 17 aatatrarac agctacaata ttccagggcc artcacttgc catttctcat aacagcgtca 113 gagagaaaga actgactgar acgtttgag atg aag aaa gtt ctc ctc ctg atc Met Lys Lys Val Leu Leu Leu Ile -15 aca gcc atc ttg gca gtg gct gtw ggt ttc cca gtc tct caa gac cag 161 Thr Ala Ile Leu Ala Val Ala Val Gly Phe Pro Val Ser Gln Asp Gln gaa cga gaa aaa aga agt atc agt gac agc gat gaa tta gct tca ggr Glu Arg Glu Lys Arg Ser Ile Ser Asp Ser Asp Glu Leu Ala Ser Gly 20 15 wtt ttt gtg ttc cct tac cca tat cca ttt cgc cca ctt cca cca att Xaa Phe Val Phe Pro Tyr Pro Tyr Pro Phe Arg Pro Leu Pro Pro Ile 30 cca ttt cca aga ttt cca tgg ttt aga cgt aan ttt cct att cca ata Pro Phe Pro Arg Phe Pro Trp Phe Arg Arg Xaa Phe Pro Ile Pro Ile 50 45 cct gaa tot gcc cct aca act ccc ctt cct agc gaa aag taaacaaraa Pro Glu Ser Ala Pro Thr Thr Pro Leu Pro Ser Glu Lys 60 ggaaaagtca crataaacct ggtcacctga aattgaaatt gagccacttc cttgaaraat caaaattoot gttaataaaa raaaaacaaa tgtaattgaa atagcacaca gcattotota <210> 18 <211> 17 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> 1..17 <223> Von Heijne matrix score 8.2 seg LLLITAILAVAVG/FP Met Lys Lys Val Leu Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val 10 5 1 Gly <210> 19 <211> 822 <212> DNA <213> Homo sapiens <220> <221> misc_feature <222> 260..464 <223> blastn

<221> misc_feature <222> 118..184

<223>	blastn						
<221>	misc_feature					•	٠.
	56113						
	blastn						
1000							
<221>	misc_feature	:					
	454485						
	blastn						
1000							
<221>	misc_feature	!					
	118545						
	blastn						
1000							
<221>	misc_feature	<b>:</b>					
<222>	65369						
	blastn				•		
1000							
<221×	misc_feature	•					
	61399						
	blastn						
12201							
<221>	misc_feature	2					
<222>	408458						
	blastn						
1220							
<221>	misc_feature	≘					
	60399						
	blastn						
<221>	misc feature	е					
	393432						
	blastn						
<221>	sig_peptide						
	346408				•		
	Von Heijne	matrix					
	· <b>y</b>			-			
<400>	. 19						
actco	tttta gcatago	agac ttcggc	gcca gcg	gccagcg	ctagtcgg	to tggtaagtgc	60
ctgat	accas attecat	tete tegegt	cttt tcc	tggtccc	aggcaaag	eg gasgnagate	120
ctcaa	acooc ctagto	atta acaatt	ccqq aqa	aaatcag	cggtctaa	tt aatteetetg	180
attte	rttgaa gcagtta	acca agaatc	ttca acc	ctttccc	acaaaagc	ta attgagtaca	240
catto	ctott gagtac	acgt tcctqt	tgat tta	caaaagg	tgcaggta	tg ageaggterg	300
aagag	taaca ttttqt	gaag ttgtaa	aaca gaa	aacctgt	tagaa at	g tgg tgg tit	357
		J. J J			Me	t Trp Trp Phe	
						-20	
cag (	aa ggc ctc a	at ttc ctt	cct tca	gcc ctt	gta att	tgg aca tct	405
Gln	In Gly Leu S	er Phe Leu	Pro Ser	Ala Leu	Val Ile	Trp Thr Ser	
	-15		-10		- 5		
act o	gct ttc ata t	tt tca tac	att act	gca gta	aca ctc	cac cat ata	453
Ala	Ala Phe Ile P	he Ser Tyr	Ile Thr	Ala Val	Thr Leu	His His Ile	
	L	5		10		15	
gac	co oct tta c	ct tat atc	agt gac	act ggt	aca gta	gct cca raa	501
Asp	Pro Ala Leu P	ro Tyr Ile	Ser Asp	Thr Gly	Thr Val	Ala Pro Xaa	
	2	:0		25		30	
aaa	rgc tta ttt g	gg gca atg	cta aat	att gcg	gca gtt	tta tgt caa	549
Lys	Cys Leu Phe G	ly Ala Met	Leu Asn	Ile Ala	Ala Val	Leu Cys Gln	
	35		40			45	
aaa	tagaaatcag ga	arataatt ca	acttaaag	g aakttc	attt catg	accaaa	602
Lvs							
ctct	tcaraa acatqt	cttt acaago	catat cto	cttgtatt	gctttcta	ca ctgttgaatt	662
		-					

gtctggcaat atttctgcag tggaaaattt gatttarmta gttcttgact gataaatatg gtaaggtggg cttttccccc tgtgtaattg gctactatgt cttactgagc caagttgtaw tttgaaataa aatgatatga gagtgacaca aaaaaaaaaa	722 782 822
<210> 20 <211> 21 <212> PRT <213> Homo sapiens	
<220> <221> SIGNAL <222> 121 <223> Von Heijne matrix score 5.5 seq SFLPSALVIWTSA/AF	
<pre>&lt;400&gt; 20 Met Trp Trp Phe Gln Gln Gly Leu Ser Phe Leu Pro Ser Ala Leu Val 1</pre>	
<210> 21 <211> 405 <212> DNA <213> Homo sapiens	
<220> <221> misc_feature <222> complement(103398) <223> blastn	
<221> sig_peptide <222> 185295 <223> Von Heijne matrix	
<pre>&lt;400&gt; 21 atcaccttct tctccatcct tstctgggcc agtccccarc ccagtccctc tcctgacctg cccagcccaa gtcagccttc agcacgcgct tttctgcaca cagatattcc aggcctacct ggcattccag gacctccgma atgatgctcc agtcccttac aagcgcttcc tggatgaggg tggc atg gtg ctg acc acc ctc ccc ttg ccc tct gcc aac agc cct gtg Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val</pre>	60 120 180 229
aac atg ccc acc act ggc ccc aac agc ctg agt tat gct agc tct gcc Asn Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala	277
ctg tcc ccc tgt ctg acc gct cca aak tcc ccc cgg ctt gct atg atg Leu Ser Pro Cys Leu Thr Ala Pro Xaa Ser Pro Arg Leu Ala Met Met	325
-5 - 1 - 5 - CCT gac aac taaatatcct tatccaaatc aataaarwra raatcctccc Pro Asp Asn	374
tccaraaggg tttctaaaaa caaaaaaaaa a	405

<210> 22 <211> 37 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> 1..37 <223> Von Heijne matrix score 5.9 seq LSYASSALSPCLT/AP <400> 22 Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val Asn 10 Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala Leu 20 Ser Pro Cys Leu Thr 35 <210> 23 <211> 496 <212> DNA <213> Homo sapiens <220> <221> misc_feature <222> 149..331 <223> blastn <221> misc_feature <222> 328..485 <223> blastn <221> misc_feature <222> complement(182..496) <223> blastn <221> sig_peptide <222> 196..240 <223> Von Heijne matrix <400> 23 60 aaaaaattgg tcccagtttt caccetgeeg cagggetgge tggggaggge ageggtttag attagccgtg gcctaggccg tttaacgggg tgacacgagc ntgcagggcc gagtccaagg 120 cccggagata ggaccaaccg tcaggaatgc gaggaatgtt tttcttcgga ctctatcgag 180 gcacacagac agace atg ggg att ctg tct aca gtg aca gcc tta aca ttt 231 Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe gcc ara gcc ctg gac ggc tgc aga aat ggc att gcc cac cct gca agt 279 Ala Xaa Ala Leu Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser gag aag cac aga ctc gag aaa tgt agg gaa ctc gag asc asc cac tcg 327 Glu Lys His Arg Leu Glu Lys Cys Arg Glu Leu Glu Xaa Xaa His Ser 20 15 gcc cca gga tca acc cas cac cga aga aaa aca acc aga aga aat tat 375 Ala Pro Gly Ser Thr Xaa His Arg Arg Lys Thr Thr Arg Arg Asn Tyr 40 35 424 tot toa goo tgaaatgaak cogggatoaa atggttgotg atcaragooo Ser Ser Ala atatttaaat tggaaaagtc aaattgasca ttattaaata aagcttgttt aatatgtctc 484 496 aaacaaaaaa aa

```
<210> 24
<211> 15
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> 1..15
<223> Von Heijne matrix
      score 5.5
      seq ILSTVTALTFAXA/LD
Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe Ala Xaa Ala
<210> 25
<211> 623
<212> DNA
<213> Homo sapiens
<220>
<221> sig_peptide
<222> 49..96
<223> Von Heijne matrix
<400> 25
aaagateeet geageeegge aggagagaag getgageett etggegte atg gag agg
                                                                       57
                                                      Met Glu Arg
                                                                       105
ctc gtc cta acc ctg tgc acc ctc ccg ctg gct gtg gcg tct gct ggc
Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala Ser Ala Gly
            -10
tgc gcc acg acg cca gct cgc aac ctg agc tgc tac cag tgc ttc aag
                                                                       153
Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln Cys Phe Lys
                        10
                                                                       201
gtc agc agc tgg acg gag tgc ccg ccc acc tgg tgc agc ccg ctg gac
Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser Pro Leu Asp
                                         30
caa gtc tgc atc tcc aac gag gtg gtc gtc tct ttt aaa tgg agt gta
                                                                       249
Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Lys Trp Ser Val
cgc gtc ctg ctc agc aaa cgc tgt gct ccc aga tgt ccc aac gac aac
                                                                       297
Arg Val Leu Leu Ser Lys Arg Cys Ala Pro Arg Cys Pro Asn Asp Asn
                                                                       345
atg aak ttc gaa tgg tcg ccg gcc ccc atg gtg caa ggc gtg atc acc
Met Xaa Phe Glu Trp Ser Pro Ala Pro Met Val Gln Gly Val Ile Thr
                             75
agg cgc tgc tgt tcc tgg gct ctc tgc aac agg gca ctg acc cca cag
                                                                       393
Arg Arg Cys Cys Ser Trp Ala Leu Cys Asn Arg Ala Leu Thr Pro Gln
gag ggg cgc tgg gcc ctg cra ggg ggg ctc ctg ctc cag gac cct tcg
                                                                       441
Glu Gly Arg Trp Ala Leu Xaa Gly Gly Leu Leu Leu Gln Asp Pro Ser
100
                     105
                                         110
agg ggc ara aaa acc tgg gtg cgg cca cag ctg ggg ctc cca ctc tgc
                                                                       489
Arg Gly Xaa Lys Thr Trp Val Arg Pro Gln Leu Gly Leu Pro Leu Cys.
                                     125
                120
                                                                       534
ctt ccc awt tcc aac ccc ctc tgc cca rgg gaa acc cag gaa gga
```

Leu Pro Xaa Ser Asn Pro Leu Cys Pro Xaa Glu Thr Gln Glu Gly 140 145 taacactgtg ggtgccccca cctgtgcatt gggaccacra cttcaccctc ttggaracaa 594 623 taaactctca tgcccccaaa aaaaaaaaa <210> 26 <211> 16 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> 1..16 <223> Von Heijne matrix score 10.1 seq LVLTLCTLPLAVA/SA <400> 26 Met Glu Arg Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala <210> 27 <211> 848 <212> DNA <213> Homo sapiens <220> <221> sig_peptide <222> 32..73 <223> Von Heijne matrix <400> 27 52 aactttgcct tgtgttttcc accctgaaag a atg ttg tgg ctg ctc ttt ttt Met Leu Trp Leu Leu Phe Phe ctg gtg act gcc att cat gct gaa ctc tgt caa cca ggt gca gaa aat 100 Leu Val Thr Ala Ile His Ala Glu Leu Cys Gln Pro Gly Ala Glu Asn 1 gct ttt aaa gtg aga ctt agt atc aga aca gct ctg gga gat aaa gca 148 Ala Phe Lys Val Arg Leu Ser Ile Arg Thr Ala Leu Gly Asp Lys Ala 15 tat gcc tgg gat acc aat gaa gaa tac ctc ttc aaa gcg atg gta gct 196 Tyr Ala Trp Asp Thr Asn Glu Glu Tyr Leu Phe Lys Ala Met Val Ala 30 ttc tcc atg aga aaa gtt ccc aac aga gaa gca aca gaa att tcc cat 244 Phe Ser Met Arg Lys Val Pro Asn Arg Glu Ala Thr Glu Ile Ser His 50 45 gtc cta ctt tgc aat gta acc cag agg gta tca ttc tgg ttt gtg gtt 292 Val Leu Leu Cys Asn Val Thr Gln Arg Val Ser Phe Trp Phe Val Val aca gac cct tca aaa aat cac acc ctt cct gct gtt gag gtg caa tca 340 Thr Asp Pro Ser Lys Asn His Thr Leu Pro Ala Val Glu Val Gln Ser gcc ata aga atg aac aag aac cgg atc aac aat gcc ttc ttt cta aat 388 Ala Ile Arg Met Asn Lys Asn Arg Ile Asn Asn Ala Phe Phe Leu Asn 100 95 gac caa act ctg gaa ttt tta aaa atc cct tcc aca ctt gca cca ccc 436

Asp Gln Thr Leu Glu Phe Leu Lys Ile Pro Ser Thr Leu Ala Pro Pro

	•																•
														120			
	~~~			110 gtg	000	ato	+ ~~	a++	115	a+ a	+++	aat	ata		ttt	4	84
Met	Asp	Pro	Ser	Val	Pro	Ile	Trp	Ile	Ile	Ile	Phe	Gly	Val	Ile	Phe	•	
			125					130			++-	+ < >	135	atc	taa	5	32
tgc	atc	atc	ata	gtt Val	gca	att	gca Nla	CLA	CEG	Tle	Len	Ser	61 v	Tle	Tro	_	,,,
Cys	TIE	140	TIE	Val	Ald	116	145	neu	Den	116	пец	150	Q ₁				
caa	cat		ara	aag	aac	aaa		cca	tct	gaa	ata		qac	gct	gaa	5	80
Gln	Arg	Xaa	Xaa	Lys	Asn	Lys	Glu	Pro	Ser	Glu	Val	Asp	Asp	Ala	Glu		
J	155			-, -		160					165	-	_				
rat	aak	tgt	gaa	aac	atg	atc	aca	att	gaa	aat	ggc	atc	CCC	tct	gat	6	28
Xaa	Xaa	Cys	Glu	Asn	Met	Ile	Thr	Ile	Glu	Asn	Gly	Ile	Pro	Ser	Asp		
170					175					180					185		
ccc	ctg	gac	atg	aag	gga	999	cat	att	aat	gat	gcc	ttc	atg	aca	gag	6	76
Pro	Leu	Asp	Met	Lys	Gly	Gly	His	Ile		Asp	Ala	Phe	Met	Thr	Glu		
				190					195					200		_	
				acc			tgaa	aggg	ctg 1	ttgt1	tctg	ct to	cctc	aaraa	a	7	27
Asp	Glu	Arg	Leu	Thr	Pro	Leu				,							
			205													_	
atta	aaaca	att 1	tgtti	tctg	tg tg	gact	gctga	a gca	atcc	tgaa	ata	ccaa	gag	caga	tcatat		787
wtt	ttgti	ttc a	acca	ttct	tc t	tttg	taata	a aat	tttt	gaat	gtg	cttg	aaa	aaaa	aaaaa		347
С																8	348
<21:<21:<21:<22:<22:<22:<22:<40	0 > 1 > S. 2 > 1 3 > V6	4 RT OMO : IGNA: 14 On Ho core eq L	L eijn 10. WLLF	e ma	AIHA,		Leu	Val	Thr	Ala	Ile	His	Ala				
<21 <21 <21	0>	5 NA rtif		l Se	_	ce											
<22	3> 0	ligo	nucl	eoti	de												
	0> 2	-															25
999	aaga	tgg	agat	agta	tt g	cctg											23
<21	0 > 3	0															

<210> 30 <211> 26 <212> DNA <213> Artificial Sequence <223> Olignucleotide

<400> 30 ctgccatgta catgatagag agattc

26

- <210> 31
- <211> 546
- <212> DNA
- <213> Homo sapiens
- <220>
- <221> promoter
- <222> 1..517
- <221> transcription start site
- <222> 518
- <221> protein_bind
- <222> 17..25
- <223> matinspector prediction
 name CMYB_01
 score 0.983
 sequence tgtcagttg
- <221> protein_bind
- <222> complement(18..27)
- <223> matinspector prediction name MYOD_Q6 score 0.961 sequence cccaactgac
- <221> protein_bind
- <222> complement (75..85)
- <223> matinspector prediction name S8_01 score 0.960 sequence aatagaattag
- <221> protein_bind
- <222> 94..104
- <223> matinspector prediction name S8_01 score 0.966 sequence aactaaattag
- <221> protein_bind
- <222> complement (129..139)
- <223> matinspector prediction name DELTAEF1_01 score 0.960 sequence gcacacctcag
- <221> protein_bind
- <222> complement (155..165)
- <223> matinspector prediction name GATA_C score 0.964 sequence agataaatcca
- <221> protein_bind

- <222> 170..178
- <223> matinspector prediction name CMYB_01 score 0.958 sequence cttcagttg
- <221> protein_bind
- <222> 176..189
- <223> matinspector prediction name GATA1_02 score 0.959 sequence ttgtagataggaca
- <221> protein_bind
- <222> 180..190
- <223> matinspector prediction
 name GATA_C
 score 0.953
 sequence agataggacat
- <221> protein_bind
- <222> 284..299
- <223> matinspector prediction name TALIALPHAE47_01 score 0.973 sequence cataacagatggtaag
- <221> protein_bind
- <222> 284..299
- <223> matinspector prediction
 name TAL1BETAE47_01
 score 0.983
 sequence cataacagatggtaag
- <221> protein_bind
- <222> 284..299
- <223> matinspector prediction
 name TAL1BETAITF2_01
 score 0.978
 sequence cataacagatggtaag
- <221> protein_bind
- <222> complement (287..296)
- <223> matinspector prediction name MYOD_Q6 score 0.954 sequence accatctgtt
- <221> protein_bind
- <222> complement (302..314)
- <223> matinspector prediction
 name GATA1_04
 score 0.953
 sequence tcaagataaagta
- <221> protein_bind
- <222> 393..405
- <223> matinspector prediction
 name IK1_01
 score 0.963
 sequence agttgggaattcc

540 546

<222> <223>	3934 mating name I score	pector pred [K2_01				·	٠.
<222>	mating name (score	in_bind 105 spector pred CREL_01 0.962 nce tgggaat		·			
<222>	mating name of score	in_bind 436 spector pre GATA1_02 0.950 nce tcagtga					•
<222>	compl matin name score	in_bind ement(478 spector pre SRY_02 0.951 nce taaaaca	diction	·			
<222>	486 matin name score	in_bind 493 spector pre E2F_02 0.957 nce tttagco					
<222>	compl matin name score	in_bind .ement(514 spector pre MZF1_01 c 0.975 ence tgaggg	ediction				
tcttg gttat gatag atcag atact	gcagt gatttg tgact ggacat ggagaa ttatc	cctgctaatt gaggtgtgct tgatagatac aaaaatgaca ttgagtagga gtcagctcag	agttgggtta ctattatttc aatctcccat ataagtacca tctggaaaac gagccttcct ttagaagcag tgggactaag	tggaactaaa tatgtggatt ggacaaaagc ctatagggaa gtggcaacgt ggagttggga	tatctatttc agggagatct aggcataaca ggagaaggga attccgttca	ttcagttgta tttttccaaa gatggtaagg agaggtcgta tgtgatttag	60 120 180 240 300 360 420 480

catcagtgat atggcaaatg tgggactaag ggtagtgatc agagggttaa aattgtgtgt

tttgttttag cgctgctggg gcatcgcctt gggtcccctc aaacagattc ccatgaatct

-14-

<210> 32 <211> 23 <212> DNA <213> Artificial Sequence

cttcat

23

24

name VMYB_02 score 0.985

<221> protein_bind
<222> 135..143

sequence tccaacggt

<223> Oligonucleotide <400> 32 gtaccaggga ctgtgaccat tgc <210> 33 <211> 24 <212> DNA <213> Artificial Sequence <223> Oligonucleotide <400> 33 ctgtgaccat tgctcccaag agag <210> 34 <211> 861 <212> DNA <213> Homo sapiens <220> <221> promoter <222> 1..806 <221> transcription start site <222> 807 <221> protein_bind <222> complement (60..70) <223> matinspector prediction name NFY_Q6 score 0.956 sequence ggaccaatcat <221> protein_bind <222> 70..77 <223> matinspector prediction name MZF1_01 score 0.962 sequence cctgggga <221> protein_bind <222> 124..132 <223> matinspector prediction name CMYB 01 score 0.994 sequence tgaccgttg <221> protein_bind <222> complement (126..134) <223> matinspector prediction

... ..

- <223> matinspector prediction
 name STAT_01
 score 0.968
 sequence ttcctggaa
- <221> protein_bind
- <222> complement (135..143)
- <223> matinspector prediction name STAT_01 score 0.951 sequence ttccaggaa
- <221> protein_bind
- <222> complement (252..259)
- <223> matinspector prediction
 name MZF1_01
 score 0.956
 sequence ttggggga
- <221> protein_bind
- <222> 357..368
- <223> matinspector prediction
 name IK2_01
 score 0.965
 sequence gaatgggatttc
- <221> protein_bind
- <222> 384..391
- <223> matinspector prediction name MZF1_01 score 0.986 sequence agaggga
- <221> protein_bind
- <222> complement (410..421)
- <223> matinspector prediction name SRY_02 score 0.955 sequence gaaaacaaaaca
- <221> protein_bind
- <222> 592..599
- <223> matinspector prediction
 name MZF1_01
 score 0.960
 sequence gaaggga
- <221> protein_bind
- <222> 618..627
- <223> matinspector prediction name MYOD_Q6 score 0.981 sequence agcatctgcc
- <221> protein bind
- <222> 632..642
- <223> matinspector prediction name DELTAEF1_01 score 0.958 seguence tcccaccttcc
- <221> protein_bind

```
<222> complement(813..823)
<223> matinspector prediction
     name S8_01
     score 0.992
     sequence gaggcaattat
<221> protein_bind
<222> complement(824..831)
<223> matinspector prediction
     name MZF1 01
     score 0.986
     sequence agaggga
<400> 34
tactataggg cacgcgtggt cgacggccgg gctgttctgg agcagagggc atgtcagtaa
                                                                   60
tgattggtcc ctggggaagg tctggctggc tccagcacag tgaggcattt aggtatetet
                                                                  120
eggtgacegt tggatteetg gaageagtag etgttetgtt tggatetggt agggacaggg
                                                                  180
ctcagagggc taggcacgag ggaaggtcag aggagaaggs aggsarggcc cagtgagarg
                                                                  240
ggagcatgcc ttcccccaac cctggcttsc ycttggymam agggcgktty tgggmacttr
                                                                  300
aaytcagggc ccaascagaa scacaggccc aktcntggct smaagcacaa tagcctgaat
                                                                  360
420
ccaaatcaag gtaacttgct cccttctgct acgggccttg gtcttggctt gtcctcaccc
                                                                  480
                                                                  540
agteggaact ecctaceact tteaggagag tggttttagg eccgtgggge tgttetgtte
caagcagtgt gagaacatgg ctggtagagg ctctagctgt gtgcggggcc tgaaggggag
                                                                  600
tgggttctcg cccaaagagc atctgcccat ttcccacctt cccttctccc accagaagct
                                                                  660
tgcctgagct gtttggacaa aaatccaaac cccacttggc tactctggcc tggcttcagc
                                                                  720
ttggaaccca atacctaggc ttacaggcca tcctgagcca ggggcctctg gaaattctct
                                                                  780
tectgatggt cetttaggtt tgggcacaaa atataattge eteteceete teccatttte
                                                                   840
                                                                   861
totottggga gcaatggtca c
<210> 35
<211> 20
<212> DNA
<213> Artificial Sequence
<220>
<223> Oligonucleotide
<400> 35
                                                                    20
ctgggatgga aggcacggta
 <210> 36
 <211> 20
 <212> DNA
 <213> Artificial Sequence
 <220>
 <223> Oligonucleotide
 <400> 36
                                                                    20
 gagaccacac agctagacaa
```

<210> 37 <211> 555 <212> DNA

<213> Homo sapiens

<220>	
<221>	promoter
<222>	1500
<221> <222>	transcription start site 501
<221>	protein_bind 191206
	matinspector prediction
	name ARNT_01 score 0.964
	sequence ggactcacgtgctgct
<221>	protein_bind
<222> <223>	193204 matinspector prediction
	name NMYC_01 score 0.965
	sequence actcacgtgctg
	protein_bind
<222> <223>	193204 matinspector prediction
	name USF_01 score 0.985
	sequence actcacgtgctg
	protein_bind
<222> <223>	complement (193204) matinspector prediction
	name USF_01 score 0.985
	sequence cagcacgtgagt
	<pre>protein_bind complement(193204)</pre>
<222>	matinspector prediction
	name NMYC_01 score 0.956
	sequence cagcacgtgagt
	protein_bind
	complement (193204) matinspector prediction
	name MYCMAX_02 score 0.972
	sequence cagcacgtgagt
<221>	protein_bind 195202
<222>	matinspector prediction
	name USF_C score 0.997
	sequence tcacgtgc
	protein_bind complement(195202)
	matinspector prediction
	name USF_C score 0.991

sequence gcacgtga

<221> protein_bind

<222> complement (210..217)

<223> matinspector prediction
 name MZF1_01
 score 0.968
 sequence catgggga

<221> protein_bind

<222> 397..410

<223> matinspector prediction
 name ELK1_02
 score 0.963
 sequence ctctccggaagcct

<221> protein_bind

<222> 400..409

<223> matinspector prediction
 name CETS1P54_01
 score 0.974
 sequence tccggaagcc

<221> protein_bind

<222> complement(460..470)

<223> matinspector prediction
 name AP1_Q4
 score 0.963
 sequence agtgactgaac

<221> protein_bind

<222> complement(460..470)

<223> matinspector prediction
 name AP1FJ_Q2
 score 0.961
 sequence agtgactgaac

<221> protein_bind

<222> 547..555

<223> matinspector prediction name PADS_C score 1.000 sequence tgtggtctc

60 ctatagggca cgcktggtcg acggcccggg ctggtctggt ctgtkgtgga gtcgggttga 120 aggacageat ttgtkacate tggtctactg cacetteeet etgeegtgea ettggeettt 180 kawaagctca gcaccggtgc ccatcacagg gccggcagca cacacatccc attactcaga 240 aggaactgac ggactcacgt gctgctccgt ccccatgagc tcagtggacc tgtctatgta 300 gagcagtcag acagtgcctg ggatagagtg agagttcagc cagtaaatcc aagtgattgt 360 catteetgte tgcattagta acteecaace tagatgtgaa aacttagtte ttteteatag gttgctctgc ccatggtccc actgcagacc caggcactct ccggaagcct ggaaatcacc 420 480 cgtgtcttct gcctgctccc gctcacatcc cacacttgtg ttcagtcact gagttacaga 540 ttttgcctcc tcaatttctc ttgtcttagt cccatcctct gttcccctgg ccagtttgtc tagctgtgtg gtctc

<210> 38

<211> 19

<212> DNA

<213> Artificial Sequence

<220> <223> Oligonucleotide	
<400> 38	10
ggccatacac ttgagtgac	19
<210> 39	
<211> 19 <212> DNA	
<213> Artificial Sequence	
<220>	
<223> Oligonucleotide	
<400> 39	
atatagacaa acgcacacc	19
<210> 40	
<211> 568	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 7471	
<221> sig_peptide	
<222> 799	
<223> Von Heijne matrix	
score 6.9	
seq LLLVPSALSLLLA/LL	
<221> polyA_signal	
<222> 537542	
<221> polyA_site <222> 554568	
<2227 334300	
<400> 40	
gggace atg tte ace age ace gge tee agt ggg ete tae aag geg eet	48
Met Phe Thr Ser Thr Gly Ser Ser Gly Leu Tyr Lys Ala Pro -30 -25 -20	
ctg tcg aag age ctt ctg ctg gtc ccc agt gcc ctc tcc ctc ctg ctc	96
Leu Ser Lys Ser Leu Leu Leu Val Pro Ser Ala Leu Ser Leu Leu Leu	
-15 -10 -5	144
gcc ctc ctc ctg cct cac tgc cag aag ccc ttt gtg tat gac ctt cac Ala Leu Leu Pro His Cys Gln Lys Pro Phe Val Tyr Asp Leu His	133
and hed hed hed pro his cys off bys pro the var by Abp and had	
gca gtc aag aac gac ttc cag att tgg agg ttg ata tgt gga aga ata	192
Ala Val Lys Asn Asp Phe Gln Ile Trp Arg Leu Ile Cys Gly Arg Ile	
20 25 30	240
att tgc ctt gat ttg aaa gat act ttc tgc agt agt ctg ctt att tat Ile Cys Leu Asp Leu Lys Asp Thr Phe Cys Ser Ser Leu Leu Ile Tyr	240
35 40 45	
aat ttt agg ata ttt gaa aga aga tat gga agc aga aaa ttt gca tcc	288
Asn Phe Arg Ile Phe Glu Arg Arg Tyr Gly Ser Arg Lys Phe Ala Ser	
50 55 60	

......

ttt ttg ctg ggt acc tgg gtt ttg tca gcc tta Phe Leu Leu Gly Thr Trp Val Leu Ser Ala Leu	ttt gac ttt ctc ctc 336 Phe Asp Phe Leu Leu
— ·	75
att gaa gct atg cag tat ttc ttt ggc atc act	
att gaa get atg eag tat the the gge ate ate	Ale Ale Com Ace Leu
Ile Glu Ala Met Gln Tyr Phe Phe Gly Ile Thr	Ala Ala Sel Asii Deu
80 85 90	95
cet tet gga tta ate ttt tgt tgt get ttt tge	tct gag act aaa ctc 432
Pro Ser Gly Leu Ile Phe Cys Cys Ala Phe Cys	Ser Glu Thr Lys Leu
100 105	110
ttc tta tca aga caa gct atg gca gag aac ttt	tcc atc taataaattt 481
tto tta toa aga caa got atg goa gag aac cee	Car Tle
Phe Leu Ser Arg Gln Ala Met Ala Glu Asn Phe	Ser Tre
115 120	tatotatato totataataa 541
aagagtagat tcatctgtat ggttgagagt aggctctgac	cacacaca egeneration
acctacatat ccaaaaaaaa aaaaaaaa	568
<210> 41	
<211> 569	
<212> DNA	•
<213> Homo sapiens	
22137 NOMO Baptems	
<220>	
<221> CDS	
<222> 168332	
<221> polyA_signal	
<222> 557562	•
<222> 337362	
· · · · · · · · · · · · · · · · · · ·	.,
<400> 41	60
aggagagata aggacataat agtattaagg gagagaaga	agacetttet eccectete 60
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga	agegeggege egagegeagg 120
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga	g agegegege egagegeagg 120 g agaagaa atg geg gae 176
aggagagata aggacataat agtattaagg gagagaaga	agegeggege egagegeagg 120
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga	g agegegege egagegeagg 120 g agaagaa atg geg gae 176
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc	g agegeggege egagegeagg 120 g agaagaa atg geg gae 176 Met Ala Asp
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc	g agegeggege egagegeagg 120 g agaagaa atg geg gae 176 Met Ala Asp 1 g aag ege atg tat tat 224
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgcttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr	aggegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 g aag ege atg tat tat 224 Lys Arg Met Tyr Tyr
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr	agegeggege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 g aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15
aggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc	aggegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 g aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272
aggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc	aggegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 g aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcgggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile	aggegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 g aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272
aggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcgggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30	agagagagagagagagagagagagagagagagagagag
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt	agagagagagagagagagagagagagagagagagagag
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu	agagagagagagagagagagagagagagagagagagag
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45	agegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag 320 Lys Gln Lys Gln Lys 50
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45	agegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag 320 Lys Gln Lys Gln Lys 50
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag	agegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag 320 Lys Gln Lys Gln Lys 50
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn	agegegege egagegeagg 120 agaagaa atg geg gae 176 Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag 320 Lys Gln Lys Gln Lys 50
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn	agegegege egagegeagg 120 agaagaa atg geg gac Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag Lys Gln Lys Gln Lys 50 gagtee aggtttecac 372
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcgggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn 55 aggaagcaga tggagccct ttcacagggg ctctgagaaa	agagagagagagagagagagagagagagagagagagag
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcgggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn 55 aggaagcaga tggagctcct ttcacagggg ctctgagaaa agcccacat cttcctaagg ggccccatgg cctgtttggg	agagagagagagagagagagagagagagagagagagag
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcgggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn 55 aggaagcaga tggagccct ttcacagggg ctctgagaaa	aggegegege egagegeagg 120 agaagaa atg geg gac 176 Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag 320 Lys Gln Lys Gln Lys 50 gagtee aggttteeae 372 a aactggagee gateteaaga 432 a ggeagggtag gteetggge 492 gettgttgtea egtaegtggt 552
agggggggtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcgggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn 55 aggaagcaga tggagctcct ttcacagggg ctctgagaaa agcccacat cttcctaagg ggccccatgg cctgtttggg	aggegegege egagegeagg 120 agaagaa atg geg gac 176 Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag 320 Lys Gln Lys Gln Lys 50 gagtee aggttteeae 372 a aactggagee gateteaaga 432 a ggeagggtag gteetggge 492 gettgttgtea egtaegtggt 552
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn 55 aggaagcaga tggagctcct ttcacagggg ctctgagaaa agccccacat cttcctaagg ggccccatgg cctgtttggg actgtgggcc gcctgcctgc tgatgtgggc tctaggcca	agagagagagagagagagagagagagagagagagagag
agggggcgtg gggccatggt ggtcttgcgg gcggggaaga tgccgcgcct tcgcctgccg cggctgtcaa ctcgctccgg gatacggcgc ccagcggggt cagaaagcaa cattgaatgc ttc tac aag gaa ttt tta agt aaa aat ttt cag Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Glr 5 10 aac aga gat tgg tac aag cgc aat ttt gcc atc Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile 20 25 30 aaa gtg gcc ctg gaa agg att tgg aac aag ctt Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu 40 45 aag agg agc aac taggagtcca ctctgaccca gccag Lys Arg Ser Asn 55 aggaagcaga tggagctcct ttcacagggg ctctgagaaa agccccacat cttcctaagg ggccccatgg cctgtttggg actgtgggcc gcctgcctgc tgatgtgggc tctaggcca	aggegegege egagegeagg 120 agaagaa atg geg gac 176 Met Ala Asp 1 aag ege atg tat tat 224 Lys Arg Met Tyr Tyr 15 acc ttc ttc atg gga 272 Thr Phe Phe Met Gly 35 aaa cag aaa caa aag 320 Lys Gln Lys Gln Lys 50 gagtee aggttteeae 372 a aactggagee gateteaaga 432 a ggeagggtag gteetggge 492 gettgttgtea egtaegtggt 552

<210> 42 <211> 895 <212> DNA

<213> Homo sapiens

<220> <221> CDS <222> 51..251

<221> sig_peptide <222> 51..110 <223> Von Heijne matrix score 5.3 seq ALIFGGFISLIGA/AF <221> polyA_signal <222> 849..854 <221> polyA_site <222> 882..895 <400> 42 56 ccgagagtgc cgggcggtcg gcgggtcagg gcagcccggg gcctgacgcc atg tcc Met Ser -20 egg aac etg ege ace geg etc att tte gge gge tte ate tee etg ate 104 Arg Asn Leu Arg Thr Ala Leu Ile Phe Gly Gly Phe Ile Ser Leu Ile - 5 -10 -15 ggc gcc gcc ttc tat ccc atc tac ttc cgg ccc cta atg aga ttg gag 152 Gly Ala Ala Phe Tyr Pro Ile Tyr Phe Arg Pro Leu Met Arg Leu Glu 10 5 gag tac aag aag gaa caa gct ata aat cgg gct gga att gtt caa gag 200 Glu Tyr Lys Lys Glu Gln Ala Ile Asn Arg Ala Gly Ile Val Gln Glu 25 20 15 gat gtg cag cca cca ggg tta aaa gtg tgg tct gat cca ttt ggc agg 248 Asp Val Gln Pro Pro Gly Leu Lys Val Trp Ser Asp Pro Phe Gly Arg 45 40 aaa tgagagggct gtcatcagct ctgattaaga aaggagattt cttcatgctt 301 tegattetge atggggtaca gecagteace teaccagaga atgaeggetg gagaagaaaa 361 ctctgtaata ccataaataa gagtgcttgt aataaaagac tgtgcacaag gattaatatt 421 tcccttctta agtatcaaaa gaactctgga acaaattata ccattaggaa ggttttcatg 481 attcagttga ttttccaaaa atgaagctat ctcacccagc tgggtttgga ggagcaatct 541 gcttattatt ctgtcgttac cacttactca agcgagctgt gatatgaata caagcaacca 601 gtgggctcgg gaaggtccgg gtctcttctg ccatcttcca gataagagat ttcagtaaaa 661 aactgccatg ctgagctgcc ttatagagct cttcgaaaat gttcgagttg ataaagctct 721 ttgaggacaa ggtacttcgt gcacctcatg ctgaagattg caccatgttg gaagataaat 781 atgaagcaag tcaaactaga tgcatacact tgtgtagaaa tcaataatca attaatagaa 841 895

<210> 43 <211> 691

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 20..613

<221> sig_peptide

<222> 20..82

<223> Von Heijne matrix
 score 10
 seq LWALAMVTRPASA/AP

<400> 43
ataccttaga ccctcagtc atg cca gtg cct gct ctg tgc ctg ctc tgg gcc
Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala

						-20					-15					
Leu	Ala	Met	Val	Thr	Arg -5	Pro	Ala	Ser	Ala	Ala 1	Pro	Met	GIY	ggc Gly 5	PIO .	100
gaa Glu	Leu	Ala	Gln	His	Glu	Glu	Leu	Thr 15	Leu	Leu	Phe	HIS	20 21	acc Thr	neu	148
Gln	Leu	Gly	Gln	Ala	Leu	Asn	Gly 30	Val	Tyr	Arg	Thr	Thr 35	GIU	gga Gly	TIP	196
ctg Leu	aca Thr 40	aaq	gcc Ala	agg Arg	aac Asn	agc Ser 45	ctg Leu	ggt Gly	ctc Leu	tat Tyr	ggc Gly 50	cgc Arg	aca Thr	ata Ile	gaa Glu	244
ctc Leu 55	cta	ggg ggg	cag Gln	gag Glu	gtc Val 60	agc Ser	cgg Arg	ggc	cgg Arg	gat Asp 65	gca Ala	gcc Ala	cag Gln	gaa Glu	ctt Leu 70	292
caa	gca Ala	agc Ser	ctg Leu	ttg Leu 75	gag	act Thr	cag Gln	atg Met	gag Glu 80	gag Glu	gat Asp	att Ile	ctg Leu	cag Gln 85	ctg Leu	340
cag Gln	gca Ala	gag Glu	gcc Ala 90	aca	gct Ala	gag Glu	gtg Val	ctg Leu 95	gjå aaa	gag Glu	gtg Val	gcc Ala	cag Gln 100	gca Ala	cag Gln	388
aag Lys	gtg Val	cta Leu 105	caa	gac Asp	agc Ser	gtg Val	cag Gln 110	caa	cta Leu	gaa Glu	gtc Val	cag Gln 115	ctg Leu	agg Arg	agc Ser	436
gcc Ala	tgg Trp 120	cta	ggc Gly	cct Pro	gcc Ala	tac Tyr 125	cga	gaa Glu	ttt Phe	gag Glu	gtc Val 130	tta Leu	aag Lys	gct Ala	cac His	484
gct Ala 135	gac	aag Lys	cag Gln	agc Ser	cac His	atc	cta Leu	tgg Trp	gcc Ala	ctc Leu 145	aca Thr	ggc Gly	cac His	gtg Val	cag Gln 150	532
caa	cag Gln	agg Arg	cgg Arg	gag Glu 155	atg Met	gtg Val	gca Ala	cag Gln	cag Gln 160	His	cgg Arg	ctg Leu	cga Arg	cag Gln 165	atc Ile	580
cag Gln	gag Glu	aga Arg	Leu	cac His	aca Thr	gcg Ala	Ala	ctc Leu 175	cca Pro	gcc	tga	atct	gcc	tgga	tggaac	633
170 175 tgaggaccaa tcatgctgca aggaacactt ccacgccccg tg							tga	ggcc	cct	gtgc	aggg	691				

<210> 44

<211> 458

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 12..416

<221> sig_peptide

<222> 12..86 <223> Von Heijne matrix score 4 seq LVVMVPLVGLIHL/GW

<221> polyA_signal <222> 425..430

<221> polyA_site <222> 445..458

<pre><400> 44 gctgaagtac t atg agc ctt cgg aac ttg tgg aga gac tac aaa gtt ttg gctgaagtac t atg agc ctt cgg aac ttg tgg aga gac tac aaa gtt ttg</pre>	50
Met Ser Leu Arg Asn Leu Trp Arg Asp Tyr Lys Val Leu	٠.
-20 - 15	
and ata ata cat ftg ddg tgg tac aga	98
gtt gtt atg gtc cct tta gtt ggg ctc ata cat tog 355 Tyr Arg Val Val Met Val Pro Leu Val Gly Leu Ile His Leu Gly Trp Tyr Arg	
Val Val Met Val Flo Bed Val Fl	246
The same at a cot and add dat date date	146
atc aaa agc agc cct gtt ttc caa ata cct ata by Asp Asp Ile Pro Ile Lys Ser Ser Pro Val Phe Gln Ile Pro Lys Asn Asp Asp Ile Pro	
The Lys Ser Ser F10 var 15 20	104
5 and ago caa ato cag	194
gag caa gat agt ctg gga ctt tcd aat ctt day Lys Ser Gln Ile Gln Glu Gln Asp Ser Leu Gly Leu Ser Asn Leu Gln Lys Ser Gln Ile Gln	
	242
the tan agt ass gas got tig dat	232
ggg aag nta gca ggc ttg caa tct tca ggt add gdd 500 Ser Ser Gly Lys Glu Ala Ala Leu Asn Gly Lys Xaa Ala Gly Leu Gln Ser Ser Gly Lys Glu Ala Ala Leu Asn 50	
45 40 45 Add Add Add Add Add Add Add Add Add Ad	290
and and and and age tog all age	230
Tan Car Dhe The Ser LVS GIU GIU Met Dyb 11811	
55 60 65	338
aag aac tgg ctt ctt gta gct ggg ata tct ttc ata ggt gac cat ctt	330
Twe len Tro Leu Leu Val Ala Gly 110 bol 2110 and 1	
	386
tot oca and cad tot qua ada coo dus	300
Gly Thr Tyr Phe Leu Gin Arg Sei Ala Dy Stin Ton	
	436
85 tot caa ago aaa caa ag agt att gaa gag tgaagtaaaa taaatatttg tot caa ago aaa caa ag agt att gaa gag tgaagtaaaa taaatatttg	450
Ser Gln Ser Lys Gln Lys Ser Ile Glu Glu	
105	458
gaattactaa aaaaaaaaaa aa	201
3440000	
<210> 45	
<210> 45 <211> 2036	
<211> 2036	
<211> 2036 <212> DNA	
<211> 2036	
<211> 2036 <212> DNA <213> Homo sapiens	
<211> 2036 <212> DNA <213> Homo sapiens <220>	
<211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS	
<211> 2036 <212> DNA <213> Homo sapiens <220>	
<211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix score 3.9</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix score 3.9</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	. 60
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	60 120
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180 240
<pre><211> 2036 <212> DNA <213> Homo sapiens </pre> <pre><220> <221> CDS <222> 2761040 </pre> <pre><221> sig_peptide <222> 276485 </pre> <pre><223> Von Heijne matrix</pre>	120 180
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180 240
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180 240 293
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180 240
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180 240 293
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180 240 293
<pre><211> 2036 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2761040 <221> sig_peptide <222> 276485 <223> Von Heijne matrix</pre>	120 180 240 293

Ile Gln Glu Ser Ile Glu Arg Leu Arg Val Ile Ala Asn Glu Ile Glu -45 -40 -35	425
aag gtc cac aga ggc tgc gtc atc gcc aat gtg gtg tcc ggc ser Thr	437
-30 -25 atg ttg gca cca ttt aca gca ggg	485
Gly Ile Leu Ser Val Ile Gly Val Met Leu Ald 116 116 -5	533
-15 ctg agc ctg agc att act gca gct ggg gta ggg ctg gga ata gca tct ctg agc ctg agc att act gca gct ggg gta ggg ctg gga ata gca tct Leu Ser Leu Ser Ile Thr Ala Ala Gly Val Gly Leu Gly Ile Ala Ser 10 15	
gcc acg gct ggg atc gcc tcc agc atc gtg gag aac aca tac aca agg Ala Thr Ala Gly Ile Ala Ser Ser Ile Val Glu Asn Thr Tyr Thr Arg	581
20 25 and act acc acc act gac caa	629
Ser Ala Glu Leu Thr Ala Ser Arg Leu III Ala III 45	677
ttg gag gca tta agg gac att ctg cat gac atc aca ccc aat gtg ctc	677
50 55 con acc aca and att acq aat gat	725
Ser Phe Ala Leu Asp Phe Asp Glu Ala III By 80	773
gtc cat aca ctc agg aga tct aaa gcc act gtt gga cgc cct ttg att yal His Thr Leu Arg Arg Ser Lys Ala Thr Val Gly Arg Pro Leu Ile 90 95	
gct tgg cga tat gta cct ata aat gtt gtt gag aca ctg aga aca cgt Pla Tro Arg Tyr Val Pro Ile Asn Val Val Glu Thr Leu Arg Thr Arg	821
100 105 at a gcc cgg aac ctg ggc aag	869
Gly Ala Pro Thr Arg He val Arg Lys val Ard 125	917
gcc act tca ggt gtc ctc gtt gtg ctg gat gta gtc aac ctt gtg caa Ala Thr Ser Gly Val Leu Val Val Leu Asp Val Val Asn Leu Val Gln 130 130 130	
gac tca ctg gac ttg cac aag ggg gaa aaa tcc gag tct gct gag ttg Asp Ser Leu Asp Leu His Lys Gly Glu Lys Ser Glu Ser Ala Glu Leu	965
145 150 150 and gag gag aat ctc aat gag ctc acc	1013
Leu Arg Gln Trp Ala Gln Glu Leu Glu Glu Abn 255 175	1060
cat atc cat cag agt cta aaa gca ggc taggcccaat tgttgcggga His Ile His Gln Ser Leu Lys Ala Gly	
180 185 agtcagggac cccaaacgga gggactggct gaagccatgg cagaagaacg tggattgtga	1120 1180
agtcagggac cccaaacgga gggactggct gatgactats business ctatgcctgt agatttcatg gacatttatt agttccccaa attaatactt trataatttc ctatgcctgt agatttcat ggacacttat cacttcccca ctttaccgca atctctaaac acaattgtg aagatttta atctcctaat cctgtcagct	1240
	1300
	1360 1420
	1420
	1540
	1600
	1660
	1720
	1780
	1840
acetcattag caattttaat tteteceegg teetgggte etgtgatete aceetgeete eaettgeett gtgatattet attacettgt gaagtaggtg atetttgtga eceacaceet eaettgeett gtgatattet attacettgt gaagtaggtg atetttgtgt tttgeagett	1900
	1960
gtgaggcatc acggaaccta ctgatgtgtg atgtctcccc tggacaccta gctttaaaat	2020 2036
ttcaaaaaa aaaaaa	2030

-26-

```
<210> 46
<211> 1276
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 443..619
<221> sig_peptide
<222> 443..589
<223> Von Heijne matrix
      score 7
      seq LICVVCLYIVCRC/GS
<221> polyA_site
<222> 1267..1276
<400> 46
gaggcactca cggcatttca ttgctacttt aattttcatt attatgggat tgattgctgt
                                                                        60
cacagctact gctgcagtag ctggagttgc tttgcattcc acagtacaaa cagcagacta
                                                                       120
tgtaaataat tggtagaaaa attctactct gctgtggaat taccaagata atatagacca
                                                                       180
gaaactaget gateaaatta atgateteea acaaactgta atgtggetag gggateatat
                                                                       240
agttagttta gaatatagaa tgcggttaca atgtgattga aatacctctg atttttgcat
                                                                       300
tactcctcat ctgtgtaatg aaacagagca tgagtgggaa aaagttaaga gatatttaaa
                                                                       360
 aggicatact agaaatttat ctttggatat tgcaaagcta aaggaacaag tatttcaagc
                                                                       420
coctcagata catotgacao ta atg coa gga act gaa gtg ott gaa gga got
                                                                       472
                          Met Pro Gly Thr Glu Val Leu Glu Gly Ala
                                          -45
 aca gac gga tta gca gct att aac ctg cta aaa tgg atc aag aca ctt
                                                                       520
 Thr Asp Gly Leu Ala Ala Ile Asn Leu Leu Lys Trp Ile Lys Thr Leu
                                                          -25
                                     -30
                 -35
 gga ggc tct gtg att tca atg att gtg ctt tta atc tgt gtt gtt tgt
                                                                       568
 Gly Gly Ser Val Ile Ser Met Ile Val Leu Leu Ile Cys Val Val Cys
                                                     -10
                                  -15
             -20
 ctt tat ata gtc tgt aga tgc gga agc cac ctc tgg aga gaa agc cac
                                                                       616
 Leu Tyr Ile Val Cys Arg Cys Gly Ser His Leu Trp Arg Glu Ser His
                             1
                                                                        669
 cac tgagagcaag caatgatagc tgtggcggtt ttgcaaaaag aaaagggaga
 His
 caagcgccca gctatagtta ccaataaagc atggtactgg tattaaaata ggcatgtgtt
                                                                        729
 ctgttccaat ggaacagaat agagaaccca gaaacaaagc caaatattta cagccaactg
                                                                        789
 atctctgaca aagcaaacaa aaacataaag tggggaaagg acaccctatt ccacaaatag
                                                                        849
 tgcagggata attggcaagc cacatgtaga aaaatgaagc tggatcctcg tctctcactt
                                                                        909
 tatacaaaaa tcaactcaaa atgggtcaaa gtcttaactc taagacctga aaccataaca
                                                                        969
 attctagaaa ataacattgg aaaaactctt ctagacattg gtttaggcaa aaagttcatg
                                                                       1029
 accaagaacc caaaagcaaa tgcaataaaa aggaagataa atagatggga cctaattaag
                                                                       1089
 ctgaaaagct tctgcatagc aaaaggaata atcagcagag caaacagaca acccacaggg
                                                                       1149
 tgggagaaaa tatttgcaag ctatgtatct gacaatggac taatatccag aatctacaag
                                                                       1209
 gaattcaaac aattagcaag aaaaaacact tgtattgtgt ttgctctgta aatcagcaaa
                                                                       1269
                                                                       1276
  aaaaaaa
```

<210> 47 <211> 747 <212> DNA <213> Homo sapiens .

<220>
<221> CDS
<222> 206..745

<400> 47 accagaagca ggtgatttcc gagctcagca atgctcagct cataatgatg tcaagcacca tggccagttt tatgaatggc ttcctgtgtc taatgaccct gacaacccat gttcactcaa gtgccaagcc aaaggaacaa ccctggttgt tgaactagca cctaaggtct tagatggtac 180 gegttgetat acagaatett tggat atg tge atc agt ggt tta tge caa att Met Cys Ile Ser Gly Leu Cys Gln Ile gtt ggc tgc gat cac cag ctg gga agc acc gtc aag gaa gat aac tgt 280 Val Gly Cys Asp His Gln Leu Gly Ser Thr Val Lys Glu Asp Asn Cys 15 ggg gtc tgc aac gga gat ggg tcc acc tgc cgg ctg gtc cga ggg cag 328 Gly Val Cys Asn Gly Asp Gly Ser Thr Cys Arg Leu Val Arg Gly Gln 35 tat aaa tcc cag ctc tcc gca acc aaa tcg gat gat act gtg gtt gca 376 Tyr Lys Ser Gln Leu Ser Ala Thr Lys Ser Asp Asp Thr Val Val Ala 50 att ccc tat gga agt aga cat att cgc ctt gtc tta aaa ggt cct gat 424 Ile Pro Tyr Gly Ser Arg His Ile Arg Leu Val Leu Lys Gly Pro Asp 70 65 cac tta tat ctg gaa acc aaa acc ctc cag ggg act aaa ggt gaa aac 472 His Leu Tyr Leu Glu Thr Lys Thr Leu Gln Gly Thr Lys Gly Glu Asn 85 80 agt ctc age tcc aca gga act ttc ctt gtg gac aat tct agt gtg gac 520 Ser Leu Ser Ser Thr Gly Thr Phe Leu Val Asp Asn Ser Ser Val Asp 100 95 ttc cag aaa ttt cca gac aaa gag ata ctg aga atg gct gga cca ctc 568 Phe Gln Lys Phe Pro Asp Lys Glu Ile Leu Arg Met Ala Gly Pro Leu 115 110 aca gca gat ttc att gtc aag att cgt aac tcg ggc tcc gct gac agt 616 Thr Ala Asp Phe Ile Val Lys Ile Arg Asn Ser Gly Ser Ala Asp Ser 130 125 aca gtc cag ttc atc ttc tat caa ccc atc atc cac cga tgg agg gag 664 Thr Val Gln Phe Ile Phe Tyr Gln Pro Ile Ile His Arg Trp Arg Glu 150 145 acg gat ttc ttt cct tgc tca gca acc tgt gga gga ggt tat cag ctg 140 712 Thr Asp Phe Phe Pro Cys Ser Ala Thr Cys Gly Gly Gly Tyr Gln Leu 165 160 747 aca tog got gag tgc tac gat ctg agg agc aac og Thr Ser Ala Glu Cys Tyr Asp Leu Arg Ser Asn 175 170

<210> 48

<211> 561

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 36..521

<221> sig_peptide

<222> 36..104

<223> Von Heijne matrix
score 7.4
seq VLLLAALPPVLLP/GA

<221> polyA_signal <222> 528..533 <221> polyA_site <222> 548..561 gacgcctctt tcagcccggg atcgccccag caggg atg ggc gac aag atc tgg 53 Met Gly Asp Lys Ile Trp -20 etg ecc tte ecc gtg etc ett etg gee get etg ect ecg gtg etg etg 101 Leu Pro Phe Pro Val Leu Leu Leu Ala Ala Leu Pro Pro Val Leu Leu -10 cct ggg gcg gcc ggc ttc aca cct tcc ctc gat agc gac ttc acc ttt 149 Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu Asp Ser Asp Phe Thr Phe acc ctt ccc gcc ggc cag aag gag tgc ttc tac cag ccc atg ccc ctg 197 Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe Tyr Gln Pro Met Pro Leu 25 20 aag gcc tcg ctg gag atc gag tac caa gtt tta gat gga gca gga tta 245 Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val Leu Asp Gly Ala Gly Leu 40 gat att gat ttc cat ctt gcc tct cca gaa ggc aaa acc tta gtt ttt 293 Asp Ile Asp Phe His Leu Ala Ser Pro Glu Gly Lys Thr Leu Val Phe 55 gaa caa aga aaa tca gat gga gtt cac act gta gag act gaa gtt ggt 341 Glu Gln Arg Lys Ser Asp Gly Val His Thr Val Glu Thr Glu Val Gly 70 gat tac atg ttc tgc ttt gac aat aca ttc agc acc att tct gag aag 389 Asp Tyr Met Phe Cys Phe Asp Asn Thr Phe Ser Thr Ile Ser Glu Lys 90 85 437 gtg att ttc ttt gaa tta atc ccg gat aat atg gga gaa cag gca caa Val Ile Phe Phe Glu Leu Ile Pro Asp Asn Met Gly Glu Gln Ala Gln 105 100 485 gaa caa gaa gat tgg aag aaa tat att act ggc aca gat ata ttg gat Glu Gln Glu Asp Trp Lys Lys Tyr Ile Thr Gly Thr Asp Ile Leu Asp 120 115 atg aaa ctg gaa gac atc ctg gtc agt atg gtc ttc taataaaata 531 Met Lys Leu Glu Asp Ile Leu Val Ser Met Val Phe

135

<210> 49

<211> 632

<212> DNA

<213> Homo sapiens

130

aaaattatta acagccaaaa aaaaaaaaaa

<220>

<221> CDS

<222> 36..395

<221> sig_peptide

<222> 36..104

<223> Von Heijne matrix score 7.4 seq VLLLAALPPVLLP/GA

<221> polyA_signal <222> 599..604

<221> polyA_site <222> 619632	•.
<pre><400> 49 gacgcctctt tcagcccggg atcgccccag caggg atg ggc gac aag atc tgg Met Gly Asp Lys Ile Trp</pre>	53
ctg ccc ttc ccc gtg ctc ctt ctg gcc gct ctg cct ccg gtg ctg c	101
cct ggg gcg gcc ggc ttc aca cct tcc ctc gat agc gac ttc acc ttt cct ggg gcg gcc ggc ttc aca cct tcc ctc gat agc gac ttc acc ttt Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu Asp Ser Asp Phe Thr Phe 10 15	149
acc ctt ccc gcc ggc cag aag gag tgc ttc tac cag ccc atg ccc ctg Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe Tyr Gln Pro Met Pro Leu 30	197
aag gcc tcg ctg gag atc gag tac caa gtt tta gat gga gca gga tta Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val Leu Asp Gly Ala Gly Leu	245
gat att gat ttc cat ctt gcc tct cca gaa ggc aaa acc tta gtt tee Asp Ile Asp Phe His Leu Ala Ser Pro Glu Gly Lys Thr Leu Val Phe	293
gaa caa aga aaa tca gat gga gtt cac acg tgt ata aga agt aaa ddo Glu Gln Arg Lys Ser Asp Gly Val His Thr Cys Ile Arg Ser Lys Asn	389
ggg cca ggc act gcg gtt cac gcc tat aat ccc agc act ttc cga ggc ggg cca ggc act gcg gtt cac gcc tat aat ccc agc act ttc cga ggc Gly Pro Gly Thr Ala Val His Ala Tyr Asn Pro Ser Thr Phe Arg Gly 95 80 85	445
80 caa gtg tagagactga agttggtgat tacatgttct gctttgacaa tacattcagc Gln Val	505
Gln Val accatttctg agaaggtgat tttctttgaa ttaatcctgg ataatatggg agaacaggca caaggacaag aagattggaa gaaatatatt actggcacag ataatattgga tatgaaactg caaggacaag aagattggaa gaaatataa taataa actggcacag attattaaca gccaaaaaaa	565
caaggacaag aagattggaa gaadtatatt actggatadag attattaaca gccaaaaaaa gaagacatcc tggtcagtat ggtcttctaa taaaataaaa	625 632
<210> 50 <211> 370	
<212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 2141	
<221> polyA_signal	
<221> polyA_site <222> 357370	
<pre><400> 50 ctgggacttc tggcctcaca atg gtt gag atg act ggg gtg tagcagtgcc</pre>	51
aagtcgagge tgtgaaagge ettecacett tactetegtg etegtgeeet eeeceattgt taggagaaagg geatgeteag gecageeat tageeeagga ggaggacaag aaacacaegg ageagacaca ageeacetea eeaaceeage caaggetgte etgaattage aaceetgaca egtgtgagea agteeaaegg acaceggaag ateeacetag teaageeeaa eeaagaetgg cagagetgee aagetgacea ettaaggege atgaggaata aacactegtt getgeatgee	111 171 231 291 351 370

attgcaaaaa aaaaaaaaa

<210> <211> <212> <213>	994 DNA		pien	s													
<220> <221> <222>	CDS																
<221: <222: <223:	35. Von	.160 Hei ore 8) ijne	mati		ſV											
<221: <222:				1													
<221: <222:																	
<400 ataa	> 51 ttgg:	ag c	tgcaa	aagc	a ga	tcgt	gaca	aga	g at Me	g ga t As	c gg p Gl:	y 01.	g aa n Ly	g aa s Ly	a aat s Asn		55
tgg Trp	Lys .	Asp	Lys '	Val	Val . -30	Asp	Leu	ьеи	TAT	-25	aga Arg	gac. Asp		-1-	-20		03
-35 act Thr	gga Gly	gtg Val	gtg Val		aat	gcc Ala	agc Ser	cta Leu	ttc Phe -10	ctg Leu	ctg Leu	ctt Leu	tca Ser	ttg Leu -5	aca Thr	1	51
gta Val	ttc Phe	agc Ser	att Ile	ata	agc Ser	gta Val	aca Thr 5	gcc Ala	tac	att Ile	gcc Ala	ttg Leu 10	gcc Ala	ctg Leu	ctc Leu	1	.99
tct Ser	Val	acc Thr	1 atc Ile	agc Ser	ttt Phe	agg Arg 20	ata	tac Tyr	aag Lys	ggt Gly	gtg Val 25	atc	caa Gln	gct Ala	atc Ile	2	47
Gln	15 aaa Lys	tca Ser	gat Asp	gaa Glu	Gly	cac	cca Pro	ttc Phe	agg Arg	gca Ala 40	tat	ctg Leu	gaa Glu	tct Ser	gaa Glu 45	2	95
30 gtt Val	gct Ala	ata Ile	tct Ser	Glu	35 gag Glu	ttg Leu	gtt Val	cag Gln	aag Lys 55	tac	agt Ser	aat Asn	tct Ser	gct Ala 60	ctt Leu	3	343
ggt Gly	cat His	gtg Val	aac Asn	50 tgc Cys	acg Thr	ata Ile	aag Lys	gaa Glu 70	ctc	agg Arg	cgc Arg	ctc Leu	ttc Phe 75	tta Leu	gtt Val	:	391
gat Asp	gat Asp	Leu	65 gtt Val	gat Asp	tct Ser	ctg Leu	aag Lys 85	ttt	gca Ala	gtg Val	ttg Leu	atg Met 90	tgg Trp	gta Val	ttt Phe	•	439
acc Thr	Tyr	80 gtt Val	ggt Gly	gcc Ala	ttg Leu	ttt Phe 100	aat	ggt Gly	ctg Le u	aca Thr	cta Leu 105	ctg Leu	att Ile	ttg Leu	gct Ala	•	487
Leu	Ile	tca Ser	ctc Leu	ttc Phe	Ser	att	cct Pro	gtt Val	att Ile	tat Tyr 120	gaa Glu	cgg	cat His	cag Gln	gca Ala 125		535
110 cag Gln		gat Asp	cat His	Tyr	Leu	gta Val	ctt Leu	gca Ala	aat Asn 135	aag Lys	aat	gtt Val	aaa Lys	gat Asp 140	gct		583
atg	gct	aaa	atc	130 caa	gca	aaa	ato	cct			aag	cgc	aaa		gaa		631

	-,
Met Ala Lys Ile Gln Ala Lys Ile Pro Gly Leu Lys Arg Lys Ala Glu	
145	691
	751
the enterty attribute attribute didaggadad dilactiget begans	811 871
terestanta atattaaata tigtaageta ctatqtatgg atttadaceg taateatate	931
the tractat statetorage cacteging ataaaaaacc tetalactic accompany	991
agatagtett geogeatett ggeaagttge agagatggtg gagetagaaa aaaaaaaaac	994
aaa	
	•
<210> 52	
<211> 412	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 271399	
<400> 52	
greatings oftogagoga togacotoot ttocaactog gcaaaccccg cttocagcag	60
	120 180
	240
ctgcgcccgg gggtcaccaa tgaacagctc tggagtgcac agaaaatcaa gcaggctatt	294
ctacatccgg acaccaatga gaagatcttc atg cca ttt aga atg tca ggt tat Met Pro Phe Arg Met Ser Gly Tyr	
1 5	2.42
att cet ttt ggg acg cca att gta agt gtt acc ttc aaa gga ttt cet	342
Ile Pro Phe Gly Thr Pro Ile Val Ser Val Thr Phe Lys Gly File Flo	
10 15 20 ttt cta aaa aat tat ttt aaa tgt cta act tta tgt tat tgc tca cgg	390
Phe Leu Lys Asn Tyr Phe Lys Cys Leu Thr Leu Cys Tyr Cys Ser Arg	
25 30 35 40	
gta ttt gac tgaattgttg att	412
Val Phe Asp	
<210> 53	
<211> 597	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 103252	
<221> sig_peptide	
<222> 103213	
<223> Von Heijne matrix	
score 3.9	
seq PGPSLRLFSGSQA/SV	
<221> polyA_site	
<222> 588597	
.400 52	
<pre><400> 53 gaaaggtcag aggaaggagc tgtgggaagc tcgcagcagg tatcggagct taagccagtg</pre>	60
gatttggggg coctgggctc cotageoggc tgoggtgtga ga arg yag cgg god	114
Met Glu Trp Ala	

-35	162
gga aag cag cgg gac ttt cag gta agg gca gct ccg ggc tgg gat cat	162
Gly Lys Gln Arg Asp Phe Gin val Arg Ara Ara 120	
	210
ttg gcc tcc ttt cct ggc cct tct ctc cgg ctg ttt tct ggg agt cag	
Leu Ala Ser Phe Pro Gly Plo Sel Leu Alg 200 1000	
• • • • • • • • • • • • • • • • • • •	252
gcg agt gtc tgt agt ctc tgc tcg ggg ttt ggg gct cag gaa Ala Ser Val Cys Ser Leu Cys Ser Gly Phe Gly Ala Gln Glu	
tagettaagg aggagggaaa cagccaattt	312
	372
	432 492
	552
	597
ctcttggtgc tgctatgtct gacctgtaat gggagaaaaa aagaa	
<210> 54	
<211> 748	
<212> DNA	
<213> Homo sapiens	
.220	
<220> <221> CDS	
<222> 2460	
<221> polyA_signal	
<222> 713718	
<221> polyA_site	
<222> 735748	
<400> 54 c aca gtt cct ctc ctc cta gag cct gcc gac cat gcc cgc ggg cgt gcc c aca gtt cct ctc ctc cta gag cct gcc gac cat gcc cgc ggg cgt gcc	49
c aca gtt cct ctc ctc cta gag cct gcc gas dat sta arg Gly Arg Ala Thr Val Pro Leu Leu Glu Pro Ala Asp His Ala Arg Gly Arg Ala	
	97
- att cac age cad tet eet gge cat gra	3,
His Val His Leu Pro Glu Ash Val Arg Scr Cim Do	
	145
cgc agg ggc aga agt ggt gca cag gta cta ccg acc gga cct gat gag	
Arg Arg Gly Arg Ser Gly Ala Gill val hear 110 1111	
	193
35 get gat the tea aag tea cat age tta	193
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu	
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 55 60 50 50 50 50 50 50 50 50	193 241
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 55 60 50 50 50 50 50 50 50 50	
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 55 60 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr	241
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 55 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 70 70 80 65	
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 55 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 70 70 80 65	241
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 55 55 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 70 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser	241
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 85 90 95	241
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 70 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 85 90 95 agc aga gga gac cat gat gac tgc cta gac ttg tgc tca gtg ctg tgt Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys	241
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 85 agc aga gga gac cat gat gac tgc cta gac ttg tgc tca gtg ctg Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 105	241 289 337
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 85 agc aga gga gac cat gat gac tgc cta gac ttg tgc tca gtg ctg Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 105 106 107 108 109 100 100 100 100 100 100	241
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 90 agc aga gga gac cat gat gac tgc cta gac ttg tgc tca gtg ctg tgt Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 100 tgg gga gaa ctg cta cgg aca ata cct gaa att cca cca aag cgt gga Trp Gly Glu Leu Leu Arg Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly	241 289 337
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50	241 289 337
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 85 agc aga gga gac cat gat gac tgc cta gac ttg tgc tca gtg ctg Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 100 tgg gga gaa ctg cta cgg aca ata cct gaa att cca cca aag cgt gga Trp Gly Glu Leu Leu Arg Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly 115	241 289 337 385
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 85 agc aga gga gac cat gat gac tgc cta gac ttg tgc tca gtg ctg Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 100 tgg gga gaa ctg cta cgg aca ata cct gaa att cca cca aag cgt gga Trp Gly Glu Leu Leu Arg Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly 115 gaa ctc aaa acg gag ctt ttg gga ctg aaa gaa aga aac cac aaa cct Glu Leu Lys Thr Glu Leu Gly Leu Lys Glu Arg Lys His Lys Pro	241 289 337 385
aaa cag gtt gag aag agt gaa gtt gat ttc tca aag tca cat agc tta Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 50 gtg aga cga ttt gag gat ctg aag ccc aag ctt tct gtt tgc aaa act Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 65 gga tca caa gtc ttt cgg tcg gag aac tgg aag gtc tgg gca gag tcg Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 85 agc aga gga gac cat gat gac tgc cta gac ttg tgc tca gtg ctg Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 100 tgg gga gaa ctg cta cgg aca ata cct gaa att cca cca aag cgt gga Trp Gly Glu Leu Leu Arg Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly 115	241 289 337 385

caa gtt tct caa cag gag gaa ctt aaa taactatgcc aagaattctg Gln Val Ser Gln Gln Glu Glu Leu Lys	480
150 145 tgaataatat aagtottaaa tatgtattto ttaatttatt gcatcaaact acttgtoott tgaataatat aagtottaaa tatgtattto ttaatttatt gtatcaaact acttgtoott	540
tgaataatat aagtottaaa tatgtattio taatotta gtggatgttt agoogatacg aagcacttag totaatgcta actgcaagag gaggtgctca gtggatgttt agoogatacg	600
	660
ttgaaattta attacggttt gattgatatt tottgatatto og attacatgc gaaatagaat aaccatttca tgaatatggt ttggaaagatg tttagtcttg aatataatgc gaaatagaat	720
aaccatttca tgaatatggt ttggaagatg tttugetotg	748
atttgtaagt ctaccaaaaa aaaaaaaa	
· ·	
<210> 55	
<211> 703	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 31231	
<2222 J1 20 -	
<221> polyA_signal	
and males of the	
<221> polyA_site	
<222> 690703	
<400> 55	54
ctctggtggc tctgctacgg cggcgcagaa atg agg cag aag cgg abb Shy Asp Met Arg Gln Lys Arg Lys Gly Asp	24
	102
ctc agc cct gct aag ctg atg atg ctg act ata gga gat gtt att aaa	
Leu Ser Pro Ala Lys Leu Met Met Bed Thi Tro	
10 15 20 gar atc gat cta aat aag	150
caa ctg att gaa gcc cac gag cag ggg aaa gac atc gat cta aat aag Gln Leu Ile Glu Ala His Glu Gln Gly Lys Asp Ile Asp Leu Asn Lys	
25 and the gar off tot got cag coc ego	198
gtg aga acc aag aca gct gcc ada tat ggc Geb Ser Ala Gln Pro Arg Val Arg Thr Lys Thr Ala Ala Lys Tyr Gly Leu Ser Ala Gln Pro Arg	
Val Arg Thr Lys Thr Ala Ala Lys 1/2 50 55	
	251
45 The state at act act acc at act act act act act a	251
ctg gtg gat atc att gct gcc gtc cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu	
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 60 67 68 68 68 68	311
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 cgccgctgcc aatttttgta tttttagtag ggatgggggt ttcaccatat tggtcaggct	311 371
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 cgccgctgcc aatttttgta tttttagtag ggatgggggt ttcaccatat tggtcaggct ggtctcgaac tcctgacctc aggtgatcaa cccaccttgg cctccctaaa tgccgggatt	311 371 431
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 cgccgctgcc aatttttgta tttttagtag ggatgggggt ttcaccatat tggtcaggct ggtctcgaac tcctgacctc aggtgatcaa cccaccttgg cctccctaaa tgccgggatt ggtctcgaac tcctgacctc cgggcctttg attttttaag gtggattttg gttgttataa acaggcatga gccaccgctc cgggcctttg attttttaag gtggattttg gttgttataa	311 371 431 491
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 cgccgctgcc aatttttgta tttttagtag ggatgggggt ttcaccatat tggtcaggct ggtctcgaac tcctgacctc aggtgatcaa cccaccttgg cctccctaaa tgccgggatt acaggcatga gccaccgctc cgggcctttg atttttaag gtggattttg gttgttataa acaggcatga gtaagagttc aagttcaacc cgtgtgtgaa agcaaaacaa tggaaaacag atggagaaag gtaagagttc aagttcaacc cgtgtgtgaa agcaaaacaa tggaaaacag	311 371 431 491 551
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 cgccgctgcc aatttttgta tttttagtag ggatgggggt ttcaccatat tggtcaggct ggtctcgaac tcctgacctc aggtgatcaa cccaccttgg cctccctaaa tgccgggatt acaggcatga gccaccgctc cgggcctttg atttttaag gtggattttg gttgttataa atggagaaag gtaagagttc aagttcaac cgtgtgtgaa agcaaaacaa tggaaaacag gattggcttc ttcaaaaggct cctcttgtag aactgcctct ttgaaatttc gaggtaatct gattggcttc ttcaaaaggct cctcttgtag ttcctaagtt aaaagcatcg cttaaccttg	311 371 431 491 551 611
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 cgccgctgcc aatttttgta tttttagtag ggatgggggt ttcaccatat tggtcaggct ggtctcgaac tcctgacctc aggtgatcaa cccaccttgg cctccctaaa tgccgggatt acaggcatga gccaccgctc cgggcctttg atttttaag gtggattttg gttgttataa atggagaaag gtaagagttc aagttcaac cgtgtgtgaa agcaaaacaa tggaaaacag gattggcttc ttcaaaaggct cctcttgtag aactgcctct ttgaaatttc gaggtaatct gattggcttc ttcaaaaggct cctcttgtag ttcctaagtt aaaagcatcg cttaaccttg	311 371 431 491 551 611 671
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu 60 65 cgccgctgcc aatttttgta tttttagtag ggatgggggt ttcaccatat tggtcaggct ggtctcgaac tcctgacctc aggtgatcaa cccaccttgg cctccctaaa tgccgggatt acaggcatga gccaccgctc cgggcctttg atttttaag gtggattttg gttgttataa acaggcatga gtaagagttc aagttcaacc cgtgtgtgaa agcaaaacaa tggaaaacag atggagaaag gtaagagttc aagttcaacc cgtgtgtgaa agcaaaacaa tggaaaacag	311 371 431 491 551 611

<210> 56 <211> 725 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 305..565

A STATE OF THE PARTY OF THE PAR

·	
<221> polyA_signal <222> 694699	••
<221> polyA_site <222> 713725	
<400> 56 ctcacggtgg tgaaggtcac agggttgcag cactcccagt agaccaggag ctccgggagg	60 120
ctcacggtgg tgaaggtcac agggttgcag cattereday gatectetgt gegecaceee cagggeegge eccaegteet etgegeacea ccetgagttg gatectetgt gegecacee cagggeegge cetectgeet	180
cagggeegge eccaegteet etgegeacea eccegagetg gattores ecteetgeet tgagttggat ecagggetag etgetgttga ecteeceact eccaegetge ecteetgeet	240
tgagttggat ccagggctag ctgctgttga cctcccatt booms gcagccatga cgcccctgct caccctgatc ctggtggtcc tcatgggctt acctctggcc gcagccatga cgcccctgct caccctgatc caggatatca ttgccatcct	300
gcagccatga cgcccctgct caccctgate ctggtggtct caggatatca ttgccatcct caggacttgg actgccacgt gtgaggacta caaatccctc caggatatca ttgccatcct	
caggeettgg actgecacgt gtgaggaeta caaaccests object gtg tee egt gea gggt atg gat gaa ett tet gag gaa gae aag ttg ace gtg tee egt gea gggt atg gat gaa ett tet gag gaa gae aag ttg ace gtg tee egt gea Met Asp Glu Leu Ser Glu Glu Asp Lys Leu Thr Val Ser Arg Ala 10	349
	397
cgg aaa ata cag cgt ttc ttg tct cag cca ttc cag gtt gct gag gtc Arg Lys Ile Gln Arg Phe Leu Ser Gln Pro Phe Gln Val Ala Glu Val	=
ttc aca ggt cat atg ggg aag ctg gta ccc ctg aag gag acc atc aaa	445
phe Thr Gly His Met Gly Lys heu val	493
gga ttc cag cag att ttg gca ggt gaa tat gac cat ctc cca gaa cag Gly Phe Gln Gln Ile Leu Ala Gly Glu Tyr Asp His Leu Pro Glu Gln	
	541
gcc ttc tat atg gtg gga ccc att gaa gaa gct gtg gca aaa gct gat Ala Phe Tyr Met Val Gly Pro Ile Glu Glu Ala Val Ala Lys Ala Asp	
65 70 73 aag ctg gct gaa gag cat tca tcg tgaggggtct ttgtcctctg tactgtctct Lys Leu Ala Glu Glu His Ser Ser	595
	655
	715
ctccttgccc ctaacccaaa aagcttcatt tttctgtgtd 55005100000000000000000000000000000000	725
aaaaaaaaaa	123
<210> 57 <211> 1705 <212> DNA <213> Homo sapiens	
<220>	
<221> CDS	
<222> 124873	
<221> sig_peptide	
<222> 124378 <223> Von Heijne matrix	
score 3.6 seq HLSVVTLAAKVKC/IP	
<221> polyA_signal	
<221> polyA_site <222> 16941705	
<400> 57 cggaggtgag gagcggcggc cccgcccggt gcgctggagg tcgaagcttc caggtagcgg cccgcagagc ctgacccagg ctctggacat cctgagccca agtcccccac actcagtgca gtg atg agt gcg gaa gtg aag gtg aca ggg cag aac cag gag caa ttt Met Ser Ala Glu Val Lys Val Thr Gly Gln Asn Gln Glu Gln Phe	60 120 160

•	•															
	-85					- B O					-75					216
cta		cta	gcc	aag	tcg	gcc	aag	ggg	gca	gcg	ctg	gcc	aca Thr	ctc	atc	210
Leu	Leu	Leu	Ala	Lys	Ser	Ala	Lys	Gly	Ala	ALA	Leu	Ala	Thr	Leu.	-55	
					-65					-00						264
cat	cag	gtg	ctg	gag	gcc	cct	ggt	gtc	tac	gtg	ttt	gga	gaa Glu	Ton.	Leu	
His	Gln	Val	Leu	Glu	Ala	Pro	Gly	Val	TYT	Val	Phe	GIY	Glu		neu	
				-50					-45					~ •		312
gac	atq	ccc	aat	gtt	aga	gag	ctg	naa	gcc	cgg	aat	ctt	cct	CCA	Ton	712
Asp	Met	Pro	Asn	Val	Arg	Glu	Leu	Xaa	Ala	Arg	Asn	Leu	FIG	Pro	Leu	
			2 5					-30					- 25			360
202	gag	act		aaq	aat	aag	ctt	cga	cac	ctc	tca	gtt	gtc Val	acc	ctg	360
Thr	Glu	Ala	Gln	Lvs	Asn	Lys	Leu	Arg	His	Leu	Ser	Val	Val	Thr	Leu	
		20					- 15					- 10				408
act	act		ata	aaq	tqt	atc	cca	tat	gca	gtg	ttg	ctg	gag	gct	CTT	408
Ala	Δla	Lvs	Val	Lvs	Cys	Ile	Pro	Tyr	Ala	Val	Leu	Leu	Glu	Ala		
	_					7				2						456
acc		cat	aat	ata	cgg	cag	ctg	gaa	gac	ctt	gtg	att	gag	gct	gtg	456
712	Leu	Ara	Asn	Val	Arg	Gln	Leu	Glu	Asp	Leu	Val	Ile	Glu	Ala	Val	
				15					20							
	act	a a c	ata		cat.	aac	tcc	ctg	gac	cag	cgc	aac	cag	cgg	ctc	504
Cat	272	200	Val	T.e.	Ara	Glv	Ser	Leu	Asp	Gln	Arg	Asn	Gln	Arg	Leu	
			2 ^					35					70			
		~~~	50	300	atc	aaa	caa	gac	atc	caq	cgc	cag	gac	ctc	agt	552
gag	get	gac	m	egc Cor	Tle	Glv	Arg	Asp	Ile	Gln	Arg	Gln	Asp	Leu	Ser	
GIu	vaı		Tyr	Ser	TIE	Gry	50				_	55				
		45			ata	cac	gaa	taa	tat	ata	aac	tqt	gag	gtc	gtg	600
gcc	att	gcc	cga	. acc	Tou	Cln	Glu	יייי דייי	CVS	Val	Gly	Cys	Glu	Val	Val	
Ala		Ala	Arg	IIII	neu	65	Olu		,	,	70	•				
	60				. ~~~	65	ata	. acc	cat	acc	aac	caa	cac	aag	gag	648
ctg	tca	ggc	att	gag	gay	Cay	V-1	Ser	· Ara	Ala	Asn	Gln	His	Lys	Glu	
	Ser	Gly	IIe	GIU	GIU	. GIII	vai	361	. Ary	85				•	90	
75					80						gac	ratt	acc	aac	ctt	696
cag	cag	ctg	ggo	cte	aag	cag	Cay	TIC	, gay	Ser	· Glu	Val	gcc Ala	Asn	Leu	
Gln	Gln	. Leu	Gly		ггх	GIL	GII	1 116	100	, 501	. 010			105	Leu	
				95				. ~a-	100	. ~~=		gea	acc	aca	tct	744
aaa	aaa	acc	att	. aaa	gtt	, acc	acc	, 90c	ם אום	. gcc	Ala	Ala	Ala	Thr	tct Ser	
Lys	: Lys	Thr			val	Tui	1111	115	z Ale	. AIC			120		Ser	
			110	)				11:	- ata	, 200	T CT A	. cca			ggc	792
cag	gac	cct	gag	g caa	a cad	CTC	act	- 9as	y CCS	720	g gat	Dro	n Ala	Pro	ggc Gly	
Glr	ı Asp			ı Glı	ı Hıs	s her	ini	. 611	n ne	MIG	9 010	139			Gly	
		125	5				130	, 		. +0-				aad	ctc	840
acc	aac	cag	g cg	c cas	g cc	ago	aag	g aa	a 900		2 000	יפפ נ יום ב	y Tays	Gly	g ctc / Leu	
Thi	: Asi	ı Glr	a Arg	g Gli	n Pro	sei	- c пA:	s ny:	S Ale	1 561	150	) )	,, _		/ Leu	
	140	)				14!				~ ~~!			aact	atic	stttcct	893
cga	a ggg	gago	e ge	c aa	g at	t tg	gtc	caa	g teg	g aa	t ty:	aaay	aacc	900	gtttcct	
Arg	g Gly	y Se:	r Al	a Ly	s Il	e Tr	o Se:	г цу	s Se	1 AS						
7 5 1	=				16	n				70	9		ctca	aaa:	accttc	953
CC	ctgg	ggat	gtg	gggt	ccc	agct	geet	gc c	tgcc	CCLL	a 99	agec	ccca	taa	ageette	1013
4	·				~~+	っったへ	ctadi	ar c	Cato	accc		aucu				1073
			200	++~+	cta i	acaa.	aaaa'	tc a	aato	cagg	L Ca	Lyce		~~		1133
		~	+~-	~++~	ata	tatt	ccat	Luc		<u> </u>	4	~~9~				1193
			+	$\alpha c = t$	cta	tece	rarr.	ca t	caca	LYLL	a cc	9454	4990	222		1253
		~~+~	~~+	atat	cta	aaac	агаа	cc c	acau	4046	L LL		9			1313
	-+~~	ata =	+~~	CCSC	++~	ctat	accc	ta c	CCCL	gacc	Lat	Lyay	Cago		-9	
-		~~~	~~~	arrt	++2	ヘヤベス	cacc	CL Q	lauaa		4 44	agco	معدد	-5-		1433
			~~~		+	tcct	аσаа	T.C. C	catc	acac	L aL	ayıc	acce			1493
~ ~		t	+~~	at co	+~~	CAAT	acta	CL G	CLLC	aacc	Ca	gage	CLUC	2~~		
				~++~	272	mato	artc	TT T	ctta	accc	. 44	Cal	CCCG		~5	
÷	~	++	~~=	2000	+++	aato	LCCL	TT T	gtga	qui	g gt	9999	ووعما	5~~	333	
ta	gatt	atat	taa	aaaa	aaa	aagg	tata	ta t	geau	atat	c ta	tata	taat	atg	acgcaga	1705
aa	taaa	tcta	tga	gaaa	tcc	aaaa	aaaa	aa a	aa							1,00

```
<210> 58
<211> 1069
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 135..206
<221> polyA_signal
<222> 850..855
<221> polyA_site
<222> 1056..1069
<400> 58
cccactccgc tetcacgact aageteteac gattaaggea egeetgeete gattgtecag
                                                                        60
cetetgecag aagaaagett ageagecage geetcagtag agacetaagg gegetgaatg
                                                                       120
agtgggaaag ggaa atg ccg acc aat tgc gct gcg gcg ggc tgt gcc act
                                                                       170
                Met Pro Thr Asn Cys Ala Ala Ala Gly Cys Ala Thr
acc tac aac aag cac att aac atc agc ttc cac agg taacctgggc
                                                                       216
Thr Tyr Asn Lys His Ile Asn Ile Ser Phe His Arg
agggagtggg ggtgacggaa actggagttc ctattgtggc tatcgcttgt gtggaaggaa
                                                                       276
caggaggatt ctgctaattc taataacttt cccagctggt agcagggaag catcgtatgt
                                                                       336
 cotttgtgtt totcaaatot goccaattgt totctgottt oggggaagot ttactcattt
                                                                       396
 tctaaaagaa atccaagtac tgtttggtca ttacccctta gtaaaaaaa gtaacaggag
                                                                       456
 gatatcgtaa ttttctactg ttttattcct ctgttagacc gggccttgac atgaatgacg
                                                                       516
 ccgtaaggga gaaagagatc ttcccaatca gcaatcaccg taaaagcctg ctgtgttccc
                                                                       576
 gttaaaatta ggaaattoto actagatgaa ttgacatggg aggcatttag atttctaata
                                                                       636
 gtcacatagt aattetgegg aggaattgag teatetttga tagecatgga attaagegat
                                                                       696
 gttaattaaa gtgcaaacga taacctttct gttcttacta gaatagagta ataaaaagaa
                                                                       756
 cctaggtttt cttttgtttg ctggaagaaa aatcaaaatt ctttagttct gtcaaaccag
                                                                        816
 aactettgaa agcaetttga acaatgeetg gaaaataaca ggtaetetgt aaatgtttae
                                                                        876
 cttctctgca agtgcctgcc acgtgcccga agaaaagaca cattaaaaag ttaagtgaca
                                                                        936
 ccagtcctga ttttatatat tttatatacc taacaacgta tatgttagta tgtagaaatt
                                                                        996
 atatecttga cettettece tacetattae gaactgtaet tetattaaaa getgecaeta
                                                                       1056
                                                                       1069
 aaaaaaaaa aaa
 <210> 59
 <211> 1084
 <212> DNA
 <213> Homo sapiens
  <220>
  <221> CDS
  <222> 135..818
  <221> polyA_signal
  <222> 909..914
  <221> polyA_site
  <222> 1071..1084
  <400> 59
  eccaeteege teteaegaet aageteteae gattaaggea egeetgeete gattgteeag
                                                                         60
  cctctgccag aagaaagctt agcagccagc gcctcagtag aggcctaagg gcgctgaatg
                                                                         120
```

agtgggaaag ggaa atg ccg acc aat tgc gct gcg gcg ggc tgt gcc act

170

•			٠.	Met	Pro	Thr	Asn	Cys	Ala	Ala	Ala	Gly	Cys	Ala	Thr	
				٦.				5					TO			218 .
acc	tac	aac	aag	cac	att	aac	atc	agc	Dhe	Cac Wie	Arg	ttt Phe	Pro	Leu	Asp	
												Phe 25				
		15	202	222	gaa	taa	a++	cgc	ctg	gtt	agg	cgc Ara	aaa	aat	ttt	266
CCL	aaa Lvg	Ara	Ara	Lvs	Glu	Trp	Val	Arg	Leu	Val	Arg	Arg	ГÀг	Asn	Phe	
																314
atg		gga	aaa	cac	act	ttt	ctt	tgt	tca	aag	cac	ttt Phe	gaa	gcc Ala	Ser	311
val	Pro	Gly	Lys	His	Thr	Phe	Leu	Cys	ser	ьуs	HIS	Phe	GIU	AIG	60	
					E 0					33		atg Met				362
tgt	ttt	gac	cta	aca	gga	Gla	Thr	Ara	Ara	Leu	Lys	Met	Asp	Ala	Val	
				<i>C</i> E					70							
663	200	att	ttt		ttt	tgt	acc	cat	ata	aag	tct	atg Met	aaa	ctc	aag	410
Pro	Thr	Ile	Phe	Asp	Phe	Cys	Thr	His	Ile	Lys	Ser	Met	-1-	Leu	гÀг	
																458
tca	agg	aat	ctt	ttg	aag	aaa	aac	aac	agt	cgt	Cer	cca	Ala	Glv	Pro	
Ser	Arg		Leu	Leu	Lys	Lys	Asn 100	Asn	Ser	Cys	261	Pro 105		1		
		95		+	220	att	act	agt	caq	caa	gta	cta	ctt	gaa	cac	506
tct	agt	T.en	Lvs	Ser	Asn	Ile	Ser	Ser	Gln	Gln	Val	Leu	Leu	Glu	His	
																554
agc		gcc	ttt	agg	aat	cct	atg	gag	gca	aaa	aag	agg	atc	att	aaa Lug	224
Ser	Tyr	Ala	Phe	Arg	Asn	Pro) Met	Glu	Ala	. LJ S		Arg	116	116	140	
					120	١				100						602
ctg	gaa	aaa	gaa	ata	gca	ago	T.AT	i aya	aya Arc	LVS	Met	Lys	Thr	Cys	cta Leu	
				3 4 5	•				136	,						
C 2 2	220	gaa	cac			act	. cga	aga	tgg	ato	aaa	gcc	atg	r tgt	ttg Leu	650
Gln	Lys	Gli	Arg	Arg	Ala	a Thi	Arg	g Arg	Tr	Ile	F Lys	a Ala		•	Leu	
																698
gta	aac	, aat	: tta	gaa	a gca	a aat	agt	gta	l tta	a cct	two	a Glv	Thi	Sei	gaa Glu	
Val	Lys			ı Glı	ı Ala	a Asi	180	r va. n	г пес	, FIG	, _L	185	;		Glu	
		175		1	- ac	c tt	2 20	c act	t cti	c cct	t tt	g gaa	gat	: ttt	aag Lys	746
cac	: acç	- T.e.	n Dro	Th:	r Ala	a Le	u Se	r Se	r Lei	ı Pro	o Le	u Gli	ı Ası	Phe	Lys	
						7 4	—				20	~				794
ato			a caa	a ga	t ca	a ca	a ga	t aa	a ac	a ct	g ct	a agt	CT	ם ממי י אפי	cta n Leu	,,,,
Ile	Le	ı Gl	u Gl	n As	p Gl	n Gl	n As	p Ly	s Th	г пе	u ne	u se:	ישני	1 War	n Leu 220	
	_				רי	Λ				41.	2					848
aaa	a ca	g ac	c aa	g ag	t ac ~ Th	c tt	c at	l la	aatt	cage	009		J J	_	atgcct	
			r Ly	2.2	_									•		
st.	satt.	catt	ctt		~	gtaa	agat	aa t	tatg	gcac	t ta	tgcc	aaaa	ttc	attattt aagaata	908 968
					- ~ t	2272	TTAC	ro a	aLLL	чьча	a 94				-	
				~~~~	-++	ttat	rtaa	aa a	taaq	L44 @	a yu	9000				1023
99	actt	aaaa	att	ttgc	taa	taaa	ttgt	gt g	tttg	aaag	g tg	aaaa	aadd	aaa	waa	

<210> 60

× **** · ·

<211> 419

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 33..290

<221> sig_peptide

<222> 33..92

<223> Von Heijne matrix score 5.4 seq WFVHSSALGLVLA/PP aatggtaggc cttcatgtga gccagttact ac atg aat ctt cat ttc cca cag <400> 60 53 Met Asn Leu His Phe Pro Gln -20 tgg ttt gtt cat tca tca gcg tta ggc ttg gtc ctg gct cca cct ttc 101 Trp Phe Val His Ser Ser Ala Leu Gly Leu Val Leu Ala Pro Pro Phe -5 tcc tct ccg ggc act gac ccc acc ttt ccg tgt att tac tgt agg cta 149 Ser Ser Pro Gly Thr Asp Pro Thr Phe Pro Cys Ile Tyr Cys Arg Leu 10 tta aat atg atc atg acc cgc ctt gca ttt tca ttc atc acc tgt tta 197 Leu Asn Met Ile Met Thr Arg Leu Ala Phe Ser Phe Ile Thr Cys Leu 30 25 tgc cca aat tta aag gaa gtt tgt ctc att ttg cca gaa aaa aat tgt 245 Cys Pro Asn Leu Lys Glu Val Cys Leu Ile Leu Pro Glu Lys Asn Cys 45 40 290 aat agt cgg cac gct gga ttt gta ggg cca gca aaa ttg cgg cag Asn Ser Arg His Ala Gly Phe Val Gly Pro Ala Lys Leu Arg Gln 65 60 tgaaactagt ttcacttcta aagcccttca tttcccacaa ggttaagctc tcgaaacccc 350 atttgatect tggttectat ttegatecte etttggaate tgaaaategg tetecatgtt 410 419 gtatgcaaa <210> 61 <211> 682 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 485..616 <221> polyA_site <222> 669..682 <400> 61 ctcctttctc attccttatc ttgcgtgttt ttaccttttt ttcataacta agtttttgag 60 gaagttagtg ttcttttcaa agaaccggtt cgaaatgtac ttttctttgc tactttttgt 120 tattttattg atcacatctt taatcttttg ttctctatac gtggcctgtt ttgatttatt 1.80 ttactattct tgctttctaa ggtaagtatt ttgttgtgta gtgctttatt tttttcatct 240 ttcttcttga ataataatga catttttagg ttataaattt tcctctggta ctcagtttgc 300 ctcattaatt ttggcagtaa gcattctcct tttattgctt tctatgtagt ctttaatttt 360 gettttaact tettetttga tetaaggatt acctacttgt taatttecaa atattatett 420 480 ggct atg tcg ccg agg ctg gag tgc agt ggt gca atc ttg gct cac tgc 529 Met Ser Pro Arg Leu Glu Cys Ser Gly Ala Ile Leu Ala His Cys 15 10 577 aac ccc cgc ctc cca ggt tca agt tat tct cct gcc tca gct act tgg Asn Pro Arg Leu Pro Gly Ser Ser Tyr Ser Pro Ala Ser Ala Thr Trp 20 gtg aga gga tcc ctt gag ccg ggg agg ttg agg ctg cag tgagccataa 626 Val Arg Gly Ser Leu Glu Pro Gly Arg Leu Arg Leu Gln ccactactct ccagcctgga taacaaaagt gagactctga ccaaaaaaaa aaaaaa 682

<210> 62 <211> 1191 <212> DNA <213> Homo sapiens				•
<220> <221> CDS <222> 54995				
<221> sig_peptide <222> 54227 <223> Von Heijne ma score 4.1 seq_LVHHCPTWG				·
<221> polyA_signal <222> 11301135				
<221> polyA_site <222> 11811191				
<400> 62 cacggctgca ctttcca	tcc cgtcgcgg	ggg ccggccgc	ta ctccggcccc ag	g atg 56 Met
cag aat gtg att aa Gln Asn Val Ile As		137 7	ca ctg gaa gtg 9	ct gag 104
-55 tac ctg acc ccg gt Tyr Leu Thr Pro Va	-,	50 	++ agg gaa aca c	igt gta 152
-40 att acc cca gaa g	- 35	~at aaa c	at cac cta qtc	ac cac 200
-25 tgt cca aca tgg c Cys Pro Thr Trp G	-20		tto aga gtg	aag gca 248
tac cta cca aca g	5	++~ a+a :	acc asa sat gtg	ccg tgc 296
tat aag cgg tgc a	•		mat daa tto daa	gct atc 344
25 att gaa gaa gat 9 Ile Glu Glu Asp A	30	+	gta gat aca tat	cac aac 392
40 aca ggt att aca g	45	++	asa dad atc aca	ctg gaa 440
aat aag gac aat Asn Lys Asp Asn	60		tra gra cta tqt	gaa gag 488
gaa gaa gat gaa Glu Glu Asp Glu		aat aca	gat atg gaa gaa	tat gaa 536
90 gag agt gga ttg Glu Ser Gly Leu	•	and act	acc cta gat aca	agg aaa 584
105	110	t gat	gct ggc ggt gaa Ala Gly Gly Gly	gat gct 632
120 att ttg caa acc	125 aga act tat	gac ctt tac	130 atc act tat gat	aaa tat 680

				140					145					Lys 150	•	728
Tyr	Gln	Thr	Pro	Arg	Leu	Trp	Leu	160	GIÀ	Tyr	Asp	Giu	165	cgg Arg	02	
Pro	Leu	Thr	gtt Val	Glu	His	Met	Tyr 175	Glu	Asp	TTE	ser	180	Asp	cat His	Vai	776
aag Lys	Lys		gtg Val	acc Thr	att Ile	gaa Glu 190	aat Asn	cat His	cct Pro	cat His	ctg Leu 195	cca Pro	cca Pro	cct Pro	ccc Pro	824
Met	185 tgt Cys	tca Ser	gtt Val	cac His	cca Pro 205	tac	agg Arg	cat His	gct Ala	gag Glu 210	gtg Val	atg Met	aag Lys	aaa Lys	atc Ile 215	872
200 att Ile	gag Glu	act Thr	gtt Val	Ala	gaa	gga Gly	Gly ggg	gga Gly	gaa Glu 225	ctt	gga Gly	gtt Val	cat His	atg Met 230	tat Tyr	920
ctt Leu	ctt Leu	att Ile	Phe	Leu	aaa Lys	ttt Phe	gta Val	caa Gln 240	gct Ala	gtc Val	att Ile	cca Pro	aca Thr 245	116	gaa Glu	968
tat Tyr	gac Asp	tac Tyr	235 aca Thr	aga Arg	cac His	ttc Phe	Thr	atg Met	taa	tgaa	gag	agca	taaa	at		1015
+ 2+	tcat	caa	tatt	taca	at t	tctc	taat	a ga	ggac	ttat	atg	Ittta	.cgc	ccat atta aaaa	tgacca aataaa aa	1075 1135 1191

```
<210> 63
<211> 1008
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 657..923
<221> sig_peptide
<222> 657..896
<223> Von Heijne matrix
     score 3.5
     seq RGLLSACAPWGDG/ST
<221> polyA_signal
<222> 957..962
<221> polyA_site
<222> 974..1008
```

ntcgnatgtg gcacaaaacc cctctgctgg ctcatgtgtg caactgagac tgcacasacc tggggtccag ctctgctgg tgggggctag agaggaagca gggagtatct gcacacaagga tgcctgcgct caggtggttg cagaagtcag tgcccaggcc ccccaacaca gtccccaaag gtccggctc cccaggcgg ggctcctcgt ttgaggggag gtgacttccc tcccagcagg ctcttggaca cagtaagctt cccaagccct gcctgagcag cctttcctcc ttgccctgtt ccccacctc cggctccagt ccagggagct cccagggaag tggtcgaccc tgccagtggc tgggccactc tgctagagcc catccgcaa gctgggggaa tcggcaaggc agcaggagca agcagaagg agcaaagac ggagaagaag aagcagaagg agcaggagca aggagagca aggagagca tggggcccg gaggagcctt gaggagccc gaggagcccc ccccacaca gctggtcat gaggagcccc gaggagcccc gaggagcccc gaggagcccc gaggagcccc gaggagcccc gaggagcccc ccccacaca agctggtcat gaggagcccc gaggagccc gaggagcccc gaggagcccc gaggagcccc gaggagcccc ccccacacac agctgggagccc agcagagagcc accccacaca gcccacacaca	tggctagctogcagagagagagagagagagagagagagagagagagaga	atgtg gcacaaaaccagctc tggggtccag cagga tgcctgcgctcaag caaag gtccggcctc gcagg ctcttggaca ctgtt ccccacctcc gtggc tgggccactc tgcgc agcatgaagg gagcc acgagccaag	ctctgctggg caggtggttg cccagcggg cagtaagctt cggctccagt tgctagagtc agcgaaagct gtgggcactt ggaaagaacc	tgggggctag cagaagtcag ggctcctcgt cccagccct ccagggagct catccgccaa ggagaagaag gatgtcggat tggggctggt	agaggaagca tgcccaggcc ttgaggggag gcctgagcag cccagggaag gctggggca aagcagaagg ctcttcaaca gaggggcccg	ccccacaca gtgacttccc cctttcctcc tggtcgaccc tcggcaaggc agcaggagca agctggtcat gaggagcctt	60 120 180 240 300 360 420 480 540 600 659
---	---	--	---	---	---	---	--

341

401

ace tet gaa eec ete aca gee tagggacagg ageggeegge ttacetggtg

ggttggggga cgtcggcagc tcgcgtacta cgccagcagg attgaggagc agagaaacag

65

Thr Ser Glu Pro Leu Thr Ala 80

cgaaacaatc tatgctgtat cgtgcctgct caatccttaa agttaac

ttgcagttgg ttgtattcag tacctgcatt tccgttggga actccacctg tacttgttat 461
tctgtggaac tttttttatt tgtagaagga gcaagaatat tgaccttact atatagcaca 521
tctgtggaac tctttttatt cgtagaagga gcaagaatat tgaccttact atatagcaca 568

	65															
211>	538															
	DNA															
213>	Hom	o sa	pien	s												
220>																
	CDS		_													
222>	151	42	6													
·221:	sig	per	tide	<u>:</u>												
	151															
223:	Vor	Hei	jne	mati	cix											
	sco	re s	5.2													
	sec	KV.	LAGI	LLGF	GLG/I	ΚV										
-221	oq <	lva s	siana	al												
<222	50!	55	10													
<221	> po	lyA_	site													
<222	> 52	75	38													
																60
-400	< 65															
<400	> 65	саа	agag	taaq	c ag	agga	taaa	caa	ctgg	aag	gaga	gcaa	gc a	caaa	gtcat	
<400 cact	> 65 gggt actt	ca a ca q	ggag cqtc	taag tgct	c ag	agga ggaa	taaa .acca	caa aga	ctgg taaa	aag gat	gaga gccc	gcaa attt	gc a tc c	caaa acca	gtcat ccaag	120 174
<400 cact catg	> 65 gggt gctt cagc	ca a ca g tc t	ggag cgtc gcct	taag tgct tttt	c ag c gt	agga ggaa ttgt	taaa acca aagc	ato	ctt	atc	acc	cag	gga	cta	gtc	120
catg caag	gctt	ca g tc t	cgtc gcct	tttt	c gt	ttgt	acca	atg Met	ctt Leu	gtc Val	acc	cag Gln	gga Gly	cta Leu -30	gtc Val	120 174
catg caag	gctt	ca g tc t	geet	tget	c gt	ttgt	aagc	atg Met	ctt Leu	gtc Val	acc Thr	cag Gln tca	gga Gly ttg	cta Leu -30	gtc Val aaa	120
catg caag	gctt	ca g tc t	geet	tget	c gt	ttgt	aagc	atg Met	ctt Leu	gtc Val	acc Thr	cag Gln tca	gga Gly ttg	cta Leu -30	gtc Val aaa	120 174
catg caag tac Tyr	gctt cagc caa Gln	ca g tc t ggt Gly	cgtc gcct tat Tyr	tttt ttg ttg Leu	gca Ala	ggaa ttgt gct Ala	aagc aat Asn	atg Met tct Ser	Ctt Leu -35 aga Arg	gtc Val ttt Phe	acc Thr gga Gly	cag Gln tca Ser	gga Gly ttg Leu -15	Cta Leu -30 CCC Pro	gtc Val aaa Lys	120 174 222
catg caag tac Tyr	gctt cagc caa Gln	ca g tc t ggt Gly	cgtc gcct tat Tyr -25	tttt ttg Leu	c gc gca	gct Ala	aat Asn	atg Met tct Ser -20	ctt Leu -35 aga Arg	gtc Val ttt Phe	acc Thr gga Gly	cag Gln tca Ser	gga Gly ttg Leu -15	Leu -30 ccc Pro	gtc Val aaa Lys	120 174
catg caag tac Tyr	gctt cagc caa Gln	ca g tc t ggt Gly	cgtc gcct tat Tyr -25	tttt ttg Leu	c gc gca	gct Ala	aat Asn	atg Met tct Ser -20	ctt Leu -35 aga Arg	gtc Val ttt Phe	acc Thr gga Gly	cag Gln tca Ser	gga Gly ttg Leu -15	Leu -30 ccc Pro	gtc Val aaa Lys	120 174 222
catg caag tac Tyr gtt Val	cago caa Gln gca Ala	ca g tc t ggt Gly ctt Leu	cgtc gcct tat Tyr -25 gct Ala	ttgct tttt ttg Leu ggt Gly	gca Ala ctc Leu	gct Ala ttg	aat Asn gga Gly	atg Met tct Ser -20 ttt Phe	ctt Leu -35 aga Arg ggc Gly	gtc Val ttt Phe ctt Leu	gga Gly gga Gly	cag Glm tca Ser aag Lys	gga ttg Leu -15 gta Val	cta Leu -30 ccc Pro tca Ser	yetc Val aaa Lys tac Tyr	120 174 222 270
catg caag tac Tyr gtt Val	caa Gln gca Ala	ca g tc t ggt Gly ctt Leu -10	cgtc gcct tat Tyr -25 gct Ala	ttgct tttt ttg Leu ggt Gly	gca Ala ctc Leu	gct Ala ttg	aat Asn gga Gly	atg Met tct Ser -20 ttt Phe	ctt Leu -35 aga Arg ggc Gly	gtc Val ttt Phe ctt Leu	acc Thr gga Gly gga Gly	cag Gln tca Ser aag Lys 1	gga Gly ttg Leu -15 gta Val	cta Leu -30 ccc Pro tca Ser	tac Tyr	120 174 222
catg caag tac Tyr gtt Val	caa Gln gca Ala	ca g tc t ggt Gly ctt Leu -10	cgtc gcct tat Tyr -25 gct Ala	ttgct tttt ttg Leu ggt Gly	gca Ala ctc Leu	gct Ala ttg	aat Asn gga Gly	atg Met tct Ser -20 ttt Phe	ctt Leu -35 aga Arg ggc Gly	gtc Val ttt Phe ctt Leu ttt	acc Thr gga Gly gga Gly	cag Gln tca Ser aag Lys 1	gga Gly ttg Leu -15 gta Val	cta Leu -30 ccc Pro tca Ser	tac Tyr	120 174 222 270
catg caag tac Tyr gtt Val ata Ile	caa Gln gca Ala gga Gly	ca gt ggt Gly ctt Leu -10 gta Val	cgtc gcct tat Tyr -25 gct Ala tgc Cys	ttg ttg Leu ggt Gly cag	gca Ala ctc Leu agt	gct Ala ttg Leu aaa Lys	aat Asn gga Gly -5 ttc	atg Met tct Ser -20 ttt Phe cat	ctt Leu -35 aga Arg ggc Gly ttt Phe	gtc Val ttt Phe ctt Leu ttt Phe	Thr gga Gly gga Gly gaa Glu	cag Gln tca Ser aag Lys 1 gat Asp	gga Gly ttg Leu -15 gta Val cag Gln	cta Leu -30 ccc Pro tca Ser ctc Leu	tac Tyr cgt Arg 20	120 174 222 270 318
catg caag tac Tyr gtt Val ata Ile 5	caa Gln gca Ala gga Gly	ca gt ggt Gly ctt Leu -10 gta Val	cgtc gcct tat Tyr -25 gct Ala tgc Cys	ttgt ttg Leu ggt Gly cag	gca Ala ctc Leu agt Ser	gct Ala ttg Leu aaa Lys	aat Asn gga Gly -5 ttc	atg Met tct Ser -20 ttt Phe cat His	ctt Leu -35 aga Arg ggc Gly ttt Phe	gtc Val ttt Phe ctt Leu ttt Phe	Thr gga Gly gga Gly gaa Glu tgc	cag Gln tca Ser aag Lys 1 gat Asp	gga Gly ttg Leu -15 gta Val cag Gln	cta Leu -30 ccc Pro tca Ser ctc Leu acc	tac Tyr cgt Arg 20	120 174 222 270 318
catg caag tac Tyr gtt Val ata Ile 5	caa Gln gca Ala gga Gly	ca gt ggt Gly ctt Leu -10 gta Val	cgtc gcct tat Tyr -25 gct Ala tgc Cys	ttgt ttg Leu ggt Gly cag	gca Ala ctc Leu agt Ser	gct Ala ttg Leu aaa Lys	aat Asn gga Gly -5 ttc	atg Met tct Ser -20 ttt Phe cat His	ctt Leu -35 aga Arg ggc Gly ttt Phe agg	gtc Val ttt Phe ctt Leu ttt Phe	Thr gga Gly gga Gly gaa Glu tgc	cag Gln tca Ser aag Lys 1 gat Asp	gga Gly ttg Leu -15 gta Val cag Gln	cta Leu -30 ccc Pro tca Ser ctc Leu acc	tac Tyr cgt Arg 20	120 174 222 270 318
tac Tyr gtt Val ata Ile 5 ggg Gly	caa Gln gca Ala gga Gly gct	ggt Gly ctt Leu -10 gta Val	tat Tyr -25 gct Ala tgc Cys	ttg ttg Leu ggt Gly cag Gln ggt	gca Ala ctc Leu agt Ser 10 cca Pro	gct Ala ttg Leu aaa Lys cag	aat Asn gga Gly -5 ttc Phe cat	atg Met tct Ser -20 ttt Phe cat His aac	ctt Leu -35 aga Arg ggc Gly ttt Phe agg Arg	gtc Val ttt Phe ctt Leu ttt Phe 15 cac	gga Gly gga Gly gaa Glu tgc Cys	cag Gln tca Ser aag Lys 1 gat Asp ctc Leu	ttg Leu -15 gta Val cag Gln ctt Leu	Leu acc Thr	tac Tyr cgt Arg 20 tgt Cys	120 174 222 270 318
tac Tyr gtt Val ata Ile 5 ggg Gly	caa Gln gca Ala gga Gly gct	ca gt ctt Leu -10 gta Val ggt	tat Tyr -25 gct Ala tgc Cys ttt	ttg ttg Leu ggt Gly cag Gln ggt Gly 25	gca Ala ctc Leu agt Ser 10 cca Pro	gct Ala ttg Leu aaa Lys cag	aat Asn gga Gly -5 ttc Phe cat	atg Met tct Ser -20 ttt Phe cat His acc	ctt Leu -35 aga Arg ggc Gly ttt Phe agg Arg	gtc Val ttt Phe ctt Leu ttt Phe 15 cac His	gga Gly gga Gly gaa Glu tgc Cys	cag Glm tca Ser aag Lys 1 gat Asp ctc Leu	ttg Leu -15 gta Val cag Gln ctt Leu	cta Leu -30 ccc Pro tca Ser ctc Leu acc Thr 35 tct	tac Tyr cgt Arg 20 tgt Cys	120 174 222 270
catg caag tac Tyr gtt Val ata Ile 5 ggg Gly gag Glu	caa Gln gca Ala gga Gly gct Ala	ca gtc t ggt Gly ctt Leu -10 gta Val ggt Gly tgc Cys	tat Tyr -25 gct Ala tgc Cys tttt Phe aaa	ttg ttg Leu ggt Gly cag Gln ggt Gly 25 ata	gca Ala ctc Leu agt Ser 10 cca Pro	gct Ala ttg Leu aaa Lys cag Gln cat	aagc aat Asn gga Gly -5 ttc Phe cat His	atg Met tct Ser -20 ttt Phe cat His aac Asn ttau	ctt Leu -35 aga Arg ggc Gly ttt Phe agg Arg 30 agt Ser	gtc Val ttt Phe ctt Leu ttt Phe 15 cac His gag	gga Gly gga Gly gaa Glu tgc Cys aag	cag Glm tca Ser aag Lys l gat Asp ctc Leu gga Gly	ttg Leu -15 gta Val cag Gln ctt Leu gac Asp	Leu -30 ccc Pro tca Ser ctc Leu acc Thr 35 tct Ser	tac Tyr cgt Arg 20 tgt Cys	120 174 222 270 318 366
catg caag tac Tyr gtt Val ata Ile 5 ggg Gly gag Glu	caa Gln gca Ala gga Gly gct Ala	ca gtc t ggt Gly ctt Leu -10 gta Val ggt Gly tgc Cys	tat Tyr -25 gct Ala tgc Cys tttt Phe aaa	ttg ttg Leu ggt Gly cag Gln ggt Gly 25 ata	gca Ala ctc Leu agt Ser 10 cca Pro	gct Ala ttg Leu aaa Lys cag Gln cat	aagc aat Asn gga Gly -5 ttc Phe cat His	atg Met tct Ser -20 ttt Phe cat His aac Asn ttau	ctt Leu -35 aga Arg ggc Gly ttt Phe agg Arg 30 agt Ser	gtc Val ttt Phe ctt Leu ttt Phe 15 cac His gag	gga Gly gga Gly gaa Glu tgc Cys aag	cag Glm tca Ser aag Lys l gat Asp ctc Leu gga Gly	ttg Leu -15 gta Val cag Gln ctt Leu gac Asp	Leu -30 ccc Pro tca Ser ctc Leu acc Thr 35 tct Ser	tac Tyr cgt Arg 20 tgt Cys	120 174 222 270 318
catg caag tac Tyr gtt Val ata Ile 5 ggg Gly gag Glu	caa Gln gca Ala gga Gly gct Ala gaa Glu	ca gt ct ggt Gly ctt Leu -10 gta Val ggt Gly tgc Cys	tat Tyr -25 Ala tgc Cys ttte Aaa Lys 40 tcc	ttg Leu ggt Gly cag Gln ggt Gly 25 ata Ile	gca Ala ctc Leu agt Ser 10 cca Pro	gct Ala ttg Leu aaa Lys cag Gln cat	aat Asn gga Gly -5 ttc Phe cat	atg Met tct Ser -20 ttt Phe cat His aac Asn ttau	ctt Leu -35 aga Arg ggc Gly ttt Phe agg Arg 30 agt Ser	gtc Val ttt Phe ctt Leu ttt Phe 15 cac His gag	gga Gly gga Gly gaa Glu tgc Cys aag	cag Glm tca Ser aag Lys l gat Asp ctc Leu gga Gly	ttg Leu -15 gta Val cag Gln ctt Leu gac Asp	Leu -30 ccc Pro tca Ser ctc Leu acc Thr 35 tct Ser	tac Tyr cgt Arg 20 tgt Cys	120 174 222 270 318 366
tac Tyr gtt Val ata Ile 5 ggg Gly gag Glu	caa Gln gca Ala Gly gct Ala gaa Glu	ca gt ct ggt Gly ctt Leu -10 gta Val ggt Gly tgc Cys	tat Tyr -25 Ala tgc Ala tgc Att Phe aaa Lys	ttg ttg Leu ggt cag Gln ggt 25 ata Ile	gca Ala ctc Leu agt Ser 10 cca Pro aag	gct Ala ttg Leu aaa Lys cag Gln cat	aagc aat Asn gga Gly -5 ttc Phe cat His gga Gly	atg Met tct Ser -20 ttt Phe cat His aac Asn ttau 45 gtga	ctt Leu -35 aga Arg ggc Gly ttt Phe agg Arg 30 agt Ser	gtc Val ttt Phe ctt Leu ttt Phe 15 cac His gag Glu	gga Gly gga Gly gaa Glu tgc Cys aag Lys	cag Gln tca Ser aag Lys l gat Asp ctc Leu gga Gly	gga Gly ttg Leu -15 gta Val cag Gln ctt Leu gac Asp 50	Leu -30 ccc Pro tca Ser ctc Leu acc Thr 35 tct Ser	tac Tyr cgt Arg 20 tgt Cys	120 174 222 270 318 366 414
tac Tyr gtt Val ata Ile 5 ggg Gly gag Glu	caa Gln gca Ala Gly gct Ala gaa Glu	ca gt ct ggt Gly ctt Leu -10 gta Val ggt Gly tgc Cys	tat Tyr -25 Ala tgc Ala tgc Att Phe aaa Lys	ttg ttg Leu ggt cag Gln ggt 25 ata Ile	gca Ala ctc Leu agt Ser 10 cca Pro aag	gct Ala ttg Leu aaa Lys cag Gln cat	aagc aat Asn gga Gly -5 ttc Phe cat His gga Gly	atg Met tct Ser -20 ttt Phe cat His aac Asn ttau 45 gtga	ctt Leu -35 aga Arg ggc Gly ttt Phe agg Arg 30 agt Ser	gtc Val ttt Phe ctt Leu ttt Phe 15 cac His gag Glu	gga Gly gga Gly gaa Glu tgc Cys aag Lys	cag Gln tca Ser aag Lys l gat Asp ctc Leu gga Gly	gga Gly ttg Leu -15 gta Val cag Gln ctt Leu gac Asp 50	Leu -30 ccc Pro tca Ser ctc Leu acc Thr 35 tct Ser	tac Tyr cgt Arg 20 tgt Cys	120 174 222 270 318 366 414

<210> 66 <211> 1747 <212> DNA <213> Homo sapiens

<220>

<221> CDS <222> 10..1062 <221> sig_peptide <222> 10..57 <223> Von Heijne matrix score 4.9 seq FIYLQAHFTLCSG/WS <221> polyA_signal <222> 1710..1715 <221> polyA_site <222> 1735..1747 geeteacea atg gtt ecc tte ate tat etg caa gee eac ttt aca ete tgt 51 Met Val Pro Phe Ile Tyr Leu Gln Ala His Phe Thr Leu Cys -10 tot ggg tgg too ago aca tac cgg gac etc cgg aag ggt gtg tat gtg 99 Ser Gly Trp Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val Tyr Val ccc tac acc cag ggc aag tgg gaa ggg gag ctg ggc acc gac ctg gta 147 Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp Leu Val 25 20 ago ato coo cat ggo coo aac gto act gtg cgt goo aac att got goo 195 Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile Ala Ala 40 35 atc act gaa toa gac aag tto tto atc aac ggo too aac tgg gaa ggo 243 Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp Glu Gly 55 291 atc ctg ggg ctg gcc tat gct gag att gcc agg cct gac gcc ccg Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala Arg Pro Asp Asp Ser Pro 70 gag cot the the gae tot one gta aag cag ace cae gth coe aac one 339 Glu Pro Phe Phe Asp Ser Leu Val Lys Gln Thr His Val Pro Asn Leu 85 387 ttc tcc ctg cag ctt tgt ggt gct ggc ttc ccc ctc aac cag tct gaa Phe Ser Leu Gln Leu Cys Gly Ala Gly Phe Pro Leu Asn Gln Ser Glu 100 gtg ctg gcc tct gtc gga ggg agc atg atc att gga ggt atc gac cac 435 Val Leu Ala Ser Val Gly Gly Ser Met Ile Ile Gly Gly Ile Asp His 120 teg etg tae aca gge agt ete tgg tat aca ece ate egg egg gag tgg 115 483 Ser Leu Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile Arg Arg Glu Trp 135 tat tat gag gtg atc att gtg cgg gtg gag atc aat gga cag gat ctg 531 Tyr Tyr Glu Val Ile Ile Val Arg Val Glu Ile Asn Gly Gln Asp Leu 150 aaa atg gac tgc aag gag tac aac tat gac aag agc att gtg gac agt 579 Lys Met Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser Ile Val Asp Ser 170 165 ggc acc acc aac ctt cgt ttg ccc aag aaa gtg ttt gaa gct gca gtc 627 Gly Thr Thr Asn Leu Arg Leu Pro Lys Lys Val Phe Glu Ala Ala Val 185 180 aaa too ato aag goa goo too too acg gag aag tto cot gac ggt tto 675 Lys Ser Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp Gly Phe 200 195 tgg cta gga gag cag ctg gtg tgc tgg caa gca ggc acc acc cct tgg 723 Trp Leu Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr Pro Trp 215 aac att ttc cca gtc atc tca ctc tac cta atg ggt gag gtt acc aac

													Val			
Gln	Ser	Phe	Arg	Ile	Thr	116	ctt Leu	PIO	0111	0	250		cgg Arg			819
Glu	Asp	Val	Ala	Thr	Ser	caa Gln	Map	Аор	O, D	265	aag Lys		gcc Ala		270	867
255 cag Gln	tca Ser	tcc Ser	acg Thr	GIA	260 act Thr	gtt Val	atg Met	gga Gly	gct Ala 280	att	atc Ile	atg Met	gag Glu	ggc Gly 285	ttc Phe	915
tac Tyr	gtt Val	gtc Val	Phe	275 gat Asp	cgg Arg	gcc Ala	cga Arg	aaa Lys 295	cga Arg	att Ile	ggc	ttt Phe	gct Ala 300	gtc Val	agc Ser	963
			290				ttc Phe	agg Arg	200	gca	aco	ata	qaa	ggc	Pro	1011
		305				gga Gly	aga Arg	a+c	. +	cta	caa	cat	tco	aca	gac Asp	1059
	320 tga												tgcg			1112
ago cago tto	tcat cagos gatag gatag gatag gaagag caac ttgg	atga gga gaag gaag attc ccaa cgtg	tgac ttcc ctg agg aaa tgc agt tgt	ttttg cctg tggc aaaaq gaag tgct attc	get g gga g ggc ggc tga tte gtg	gatga caca agcad tggca tctgd aact tttt gtac atgc	acctoractoractoractoractoractoractoracto	ag gg tg gg cg c t gc t tt	tggttacctgaatgaatgaaga	cact cagg actc actt agg agag	t tto ac g ac t gt a c t	ggtca ccac tgta gtca cacc ggca aagc	acaa caaa cctg cctc attc tcac ttgt	gtag tgcc tagg aaa ctt acg	etgegee egggeag gagaca etetgee gagacag ettaagt taaatte eaggtta eetgetg actgtat	1172 1232 1292 1352 1412 1472 1532 1592 1652 1712 1747

<210> 67

<211> 1686

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 78..491

<221> sig_peptide

<222> 78..218

<223> Von Heijne matrix score 5.8 seq LMCFGALIGLCAC/IC

<221> polyA_signal

<222> 1652..1657

<221> polyA_site

<222> 1673..1686

<400> 67
ggtatagccc accagaaagg acagagtcat ttgatgtggt cacaaaatgt gtgagtttca 60
ggtatagccc accagaaagg acagagtcat ttgatgtggt cacaaaatgt gtgagtttca 110
cactaactga gcagttc atg gag aaa ttt gtt gat ccc gga aac cac aat 110
Met Glu Lys Phe Val Asp Pro Gly Asn His Asn

						_	45				_	40				
			gat	a+ a	at t			tat	ctt	tgg	cgt	tgc	cag	ttc	ctt	158
agc	a aa	att	gat Asp	Lou	Len	Ara	Thr	Tvr	Leu	Trp	Arg	Cys	Gln	Phe	Leu	
	-35		gtg	agt	tta	~~+	tta	atq	tqc	ttt	ggg	gct	ttg	atc	gga	206
tta	CCT	חשם	gtg Val	Ser	Len	Glv	Leu	Met	Cvs	Phe	Gly	Ala	Leu	Ile	Gly	
-20	L	~~+	tgc	a++		cga	agc	tta	tat	ccc	acc	att	gcc	acg	ggc	254
CTT	tgt	212	tgc Cys	Tle	CVS	Ara	Ser	Leu	Tyr	Pro	Thr	Ile	Ala	Thr	Gly	
																202
		cat	ctc	ctt	gca	aat	ctq	tgt	aca	ctg	ggc	tca	gta	agt	tgt	302
att	TOU	Udc	ctc Leu	Len	Ala	Glv	Leu	Cys	Thr	Leu	Gly	Ser	Val	Ser	Cys	
																250
			gga	att	gaa	cta	ctc	cac	cag	aaa	cta	gag	ctc	cct	gac	350
tat	3703	712	gga Gly	Tle	Glu	Leu	Leu	His	Gln	Lys	Leu	Glu	Leu	Pro	Asp	
																200
	_	tcc	ggt	gaa	ttt	~~~	tgg	tcc	ttc	tgc	ctt	gct	tgt	gtc	tct	398
aat	y ca	Cer	ggt	Glu	Phe	Gly	Trp	Ser	Phe	Cys	Leu	Ala	Cys	Val	ser	
																446
45 ~ct	ccc	tta	cag	ttc		qct	tct	gct	ctc	ttc	atc	tgg	gct	gct	cac	440
712	Pro	Leu	cag Gln	Phe	Met	Ala	Ser	Ala	Leu	Phe	Ile	Trp	Ala	Ala	HIS	
																491
acc	aac	cac	g aga	gag	tac	acc	tta	atg	aag	gca	tat	. cgt	gtg	gca		471
Thr	Asn	Arc	, aga , Arg	Glu	Tyr	Thr	Leu	. Met	Lys	: Ala	Туг	Arg	\ val	. Ala	•	
																551
tαa	gcaa	gaa	actg	cctg	ct t	taca	atto	jc ca	tttt	tatt	: ttt	ttaa	laat	data	ctgata tatgaa	611
																671
																731
																791
																851
																911
																971
																1031
																1091
																1151
																1211
																1271
																1331
																1391
																1451
																1511
					~~+		2222	T - L	aauu							1686
tg	tctt	tttt	act aaa	aaat	aaa	atta	aaaa	icg a	aaag	ayac						

```
<210> 68
<211> 542
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 69..371

<221> sig_peptide
<222> 69..287
```

<223> Von Heijne matrix score 4

seq AVGFLFWVIVLTS/WI

<221> polyA_signal <222> 510515	,
<221> polyA_site <222> 530542	
<400> 68 tgttacttag ggtcaaggct tgggtcttgc cccgcaaacc cttgggacga cccggccca gcgcagct atg aac ctg gag cga gtg tcc aat gag gag aaa ttg aac ctg Met Asn Leu Glu Arg Val Ser Asn Glu Glu Lys Leu Asn Leu -70 -65	60 110
tgc cgg aag tac tac ctg ggg ggg ttt gct ttc ttg cct ttt ctc tgg Cys Arg Lys Tyr Tyr Leu Gly Gly Phe Ala Phe Leu Pro Phe Leu Trp -55 -50 -45	158
ttg gtc aac atc ttc tgg ttc tac cga gag gcc ttc ctt gtc cca gcc Leu Val Asn Ile Phe Trp Phe Tyr Arg Glu Ala Phe Leu Val Pro Ala -40 -35 -30	206
tac aca gaa cag agc caa atc aaa ggc tat gtc tgg cgc tca gct gtg Tyr Thr Glu Gln Ser Gln Ile Lys Gly Tyr Val Trp Arg Ser Ala Val	254
ggc ttc ctc ttc tgg gtg ata gtg ctc acc tcc tgg atc acc atc ttc Gly Phe Leu Phe Trp Val Ile Val Leu Thr Ser Trp Ile Thr Ile Phe	302
cag atc tac cgg ccc cgc tgg ggt gcc ctt ggg gac tac ctc tcc ttc Gln Ile Tyr Arg Pro Arg Trp Gly Ala Leu Gly Asp Tyr Leu Ser Phe	350
acc ata ccc ctg ggc acc ccc tgacaacttc tgcacatact ggggccctgc Thr Ile Pro Leu Gly Thr Pro	401
ttattetece aggacagget cettaaagea gaggageetg teetgggage ceetteteaa acteetaaga ettgttetea tgteecaegt tetetgetga cateeceaa taaaggacee taacttteaa aaaaaaaaa a <210 > 69 <211 > 1174 <212 > DNA <213 > Homo sapiens <220 > <221 > CDS <222 > 2757	461 521 542
<pre><221> sig_peptide <222> 2205 <223> Von Heijne matrix</pre>	
<pre><400> 69 g atg cct gag ggc ccc gag ctg cac ctg gcc agc cag ttt gtg aat gag Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu -65 -60 -55</pre>	49
gcc tgc agg gcg ctg gtg ttc ggc ggc tgc gtg gag aag tcc tct gtc Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -50 -45 -40	97
age age age cot dag gtg coc tit gag age agt god tac cgc atc toa	145

	2 5		Pro			-30					-25				•	
gct		gcc	cgc	ggc	aag	gag	ctg	cgc	ctg	ata	ctg	agc	cct	ctg	ect Pro	193
Ala	Ser	Ala	Arg	Gly	Lys	Glu	Leu	Arg	ьeu	-10	Беп	SEL	FIO	Dea	-5	
	gcc	cag	cct	caa	cag	gag	cca	ctg	gcc	ctg	gtc	ttc	cgc	ttc	ggc	241
Gly	Ala	Gln	Pro	Gln	Gln	Glu	Pro	· Leu 5	Ala	ьeu	Val	₽11¢	10	1110	017	
atg	tcc	ggc	tct	ttt	cag	ctg	gtg	ccc	cgc	gag	gag	ctg	cca	cgc	cat Hig	289
Met	Ser	Gly 15	Ser	Phe	Gln	Leu	Val 20	Pro	Arg	GIU	GIU	25	PIO	AT 9	1125	
acc	cac		cgc	ttt	tac	acg	acc	ccg	cct	ggc	ccc	cgg	ctc	gcc	cta	337
Ala	His	Leu	Arg	Phe	Tyr	Thr	Ala	Pro	Pro	GIY	40	Arg	nea	AIG	Dea	
tat	++-	ata	gac	atc	cgc	cgg	ttc	ggc	cgc	tgg	gac	ctt	a aa	gga	aag	385
Cys	Phe	Val	Asp	Ile	Arg	Arg	Phe	Gly	Arg	Trp	Asp	Leu	Gly	Gly	Lys 60	
4 =					50					22					00	433
tgg	cag	ccg	ggc Gly	cgc	999	Dro	Cvs	Val	Leu	Gln	Glu	Tyr	Gln	Gln	Phe	
				65					70					, _		
agg	gag	aat	gtg	cta	cga	aac	cta	gcg	gat	aag	gcc	ttt	gac	cgg	ccc	481
Arg	Glu	Asn	Val	Leu	Arg	Asn	Leu	A1a 85	Asp	гÀг	Ala	Pne	90	Arg	710	
atc	tgc	gag	acc	ctc	ctg	gac	cag	agg	ttc	ttc	aat	ggc	att	ggc	aac	529
Ile	Cys	Glu	Ala	Leu	Leu	Asp	100	Arg	Pne	Pne	ASII	105	110	O. J		r -
tat	ctg	cgg	gca	gag	atc	ctg	tac	cgg	ctg	aag	atc	CCC	CCC	Dho	gag Glu	577
Tyr	Leu	Arg	Ala	Glu	Ile	Leu 115	Tyr	Arg	Leu	тλг	120	PIO	FIO	1110	014	
aaq	~~~	-	tcg	gtc	ctg	gag	gcc	ctg	cag	cag	cac	agg	ccg	agc	ccg	625
Lys	Ala	Arg	Ser	Val	Leu	Glu	Ala	. Leu	Gln	GID	nıs	Arg	Pro	Ser	Pro 140	
125					130			3.00		135		сас	aat	tca		673
gag	cto	acc	ctg	ago	cag	Lvs	Tle	Arc	Thr	Lys	Leu	Gln	Asn	Ser	gac	
				145					150	1				155		
cto	cto	gag	g cta	tgt	cac	tca	gte	ccc	aag	gaa	gtg	gto	cag	ttg	ggt	721
Let	Lei	Gli	Lev	Сув	His	Ser	· Val	. Pro) LAE	Glu	ı Val	. Val	. GII	пес	Gly	
			160)				165)				1/0	,		767
gag	ggc	aaa	a gat	ggo	ago	aac	CEC Let	Cve	: Dhe	: agc	Lvs	i tys	ccgc	.904		
		17	5				180)								
acc	ecta	raac	actt	atco	cc c	tctg	gaco	t ga	attca	ccga	a ttt	ggaa	igtt	tgta	gcccta	827
~~+	- ~ = + :	ctc	aato	rcact	ag c	racto	cctca	ac tt	tatca	aatag	g tgt	ttcc	agg	cuge	gegeag	887 947
+ ~	act ca	tac	ctat	aata	cc c	acad	cttco	ad da	aggco	gagt	: gg	grgg	ge ce	acci	gaggee	1007
			~ 2 ~ ~	-at-ca	ta c	TOCAR	acato	ra to	aaaa	ccca	ז בכי	ccad	caa	aace	gcaaaaa	1067
ati	tage	agg	tgt	gtg	geg 9	gcad	cctgi	ta gi	actte	racci	. cc	ratice outsi	tac.	catt	gcagga gcactc	1127
aaa	atcg	cttg	aac	cagg	ag 9	aaaci	ccai	to to	caaaa	aaaa	a aaa	aaaa	3 3		-	1174
cag	gcct!	399C	aace	بمعم	, ua 6											

<210> 70

<211> 1285

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 2..1051

<221> sig_peptide <222> 2..205

<223> Von Heijne matrix score 7.3 seq LRLILSPLPGAQP/QQ

<221> polyA_signal <222> 1248..1253

<221> polyA_site <222> 1272..1285

<400> 70 g atg cct gag ggc ccc gag ctg cac ctg gcc agc cag ttt gtg aat gag 49 Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu -55 -60 gcc tgc agg gcg ctg gtg ttc ggc ggc tgc gtg gag aag tcc tct gtc 97 Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -45 age ege aac eet gag gtg eee ttt gag age agt gee tae ege ate tea 145 Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser -25 -30 gct tca gcc cgc ggc aag gag ctg cgc ctg ata ctg agc cct ctg cct 193 Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro -10 -15 ggg gcc cag ccc caa cag gag cca ctg gcc ctg gtc ttc cgc ttc ggc 241 Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly 1 atg tcc ggc tct ttt cag ctg gtg ccc cgc gag gag ctg cca cgc cat 289 Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His 20 ged cad etg ege tit tad acg ged eeg eet ggd eec egg etc ged eta 337 Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu 35 tgt ttc gtg gac atc cgc cgg ttc ggc cgc tgg gac ctt ggg gga aag 385 Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys 55 50 45 tgg cag ccg ggc cgc ggg ccc tgt gtc ttg cag gag tac cag cag ttc 433 Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe 65 481 agg ctg aag atc ccc ccc ttt gag aag gcc cgc tcg gtc ctg gag gcc Arg Leu Lys Ile Pro Pro Phe Glu Lys Ala Arg Ser Val Leu Glu Ala 85 80 529 ctg cag cag cac agg ccg agc ccg gag ctg acc ctg agc cag aag ata Leu Gln Gln His Arg Pro Ser Pro Glu Leu Thr Leu Ser Gln Lys Ile 577 agg acc aag ctg cag aat cca gac ctg ctg gag cta tgt cac tca gtg Arg Thr Lys Leu Gln Asn Pro Asp Leu Leu Glu Leu Cys His Ser Val 120 115 625 ccc aag gaa gtg gac cag ttg ggg ggc agg ggc tac ggg tca gag agc Pro Lys Glu Val Asp Gln Leu Gly Gly Arg Gly Tyr Gly Ser Glu Ser 135 130 673 ggg gag gag gac ttt gct gcc ttt cga gcc tgg ctg cgc tgc tat ggc Gly Glu Glu Asp Phe Ala Ala Phe Arg Ala Trp Leu Arg Cys Tyr Gly 150 atg cca ggc atg agc tcc ctg cag gac cgg cat ggc cgt acc atc tgg 721 Met Pro Gly Met Ser Ser Leu Gln Asp Arg His Gly Arg Thr Ile Trp 165 769 ttc cag ggg gat cct gga ccg ttg gca ccc aaa ggg cgc aag tcc cgc Phe Gln Gly Asp Pro Gly Pro Leu Ala Pro Lys Gly Arg Lys Ser Arg 185 180 aaa aag aaa too aag goo aca cag otg agt oot gag gac aga gtg gag 817 Lys Lys Lys Ser Lys Ala Thr Gln Leu Ser Pro Glu Asp Arg Val Glu 195

	0.65
gac get ttg cet cea age aag gee cet tee aag aca ega agg gea aag	865
Ann Ala Leu Pro Pro Ser Lys Ala Plo Ser Lys III 115	
	913
205 210 210 aga act gca acc cag cgg cct gag ggg acc agc aga gac ctt cct aag agg act gca acc cag cgg cct gag ggg acc agc	
Arg Asp Leu Pro Lys Arg Thr Ala Thi Sin Arg 120	
225	961
ctc cag cag gac cca gaa get eee ata geg cee Lys Lys Gly Arg Arg Leu Gln Gln Asp Pro Glu Ala Pro Thr Val Pro Lys Lys Gly Arg Arg	
240 tat aga cac tag aga cac agg agg gtc aag	1009
aag ggg cga cag gca gcc tct ggc cac tgg dag Pro Arg Lys Val Lys Lys Gly Arg Gln Ala Ala Ser Gly His Cys Arg Pro Arg Lys Val Lys	
655 ADV	1051
the day con day god acc toa god tot	1051
Ala han The Pro Ser Leu Giu Pro Giu Giy ini Dai ini	
	1111
270 275 tagcaggagg ctctccttgc ttgcactcac cctttcttat tgtcttgccc tgcatctggg	1171
	1231
	1285
ctgcacaact ctcatggttt taattgtace couldered aaaaaaaaaa aaaa aaaa aaaatgctgc atttttaata aactgataca tttgaactcc aaaaaaaaaa	
<210> 71	
<211> 1398	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS <222> 21171	
<222> 211/1	
<221> sig_peptide	
<222> 2205	
<223> Von Heijne matrix	
score 7.3	
seq LRLILSPLPGAQP/QQ	
<221> polyA_signal	
<222> 13681373	
<221> polyA_site	
<222> 13861398	
<400> 71	49
and of a acc age cad till gly age 343	49
Wat Dag Clu Cly pro Gill Leu his Deu hia ber die	
	97
gcc tgc agg gcg ctg gtg ttc ggc ggc tgc gtg gag aag tcc tct gtc	•
Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Gla Single	
	145
ago cgo aac cot gag gtg coo ttt gag ago agt goo tac cgc ato toa	
Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Arg	
	193
gct tca gcc cgc ggc aag gag ctg cgc ctg ata ctg agc cct ctg cct gct tca gcc cgc ggc aag gag ctg cgc ctg ata ctg agc cct ctg cct	
Ala Ser Ala Arg Gly Lys Giu Leu Arg Leu ile beu 501 505 -5	
-20 and god ctd god ctd gtd ttd cgc ttd ggc	241
ggg gcc cag ccc caa cag gag cca ctg gcc gcc	
ata con cac gag gag ctg cca cgc cat	289
atg tcc ggc tct ttt cag ctg gtg tcc cgc ggg ggg Leu Pro Arg His Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His	
Met ser Gly ser File din 200 25	

•																
gcc	cac	ctg	cgc	ttt	tac	acg	gcc	ccg	cct	ggc	CCC	cgg	ctc	gcc	cta	337
	30		Arg			35					40					
tat	ttc	gtg	gac	atc	cgc	cgg	ttc	ggc	cgc	tgg	gac	ctt	999	gga	aag	385
Cys	Phe	Val	Asp	Ile	Arg	Arg	Phe	Gly	Arg	Trp 55	Asp	Leu	Gly	Gly	Lys 60	
45					50.						~~~	+ > 0	C 3 C	CaG		433
tgg	cag	ccg	ggc	cgc	aaa	ccc	tgt	gtc	ttg	cag	gag	m	Cay	Cla	Dhe	433
Trp	Gln	Pro	Gly	Arg 65	GIY	Pro	Cys	vai	ьеu 70	GIN.	GIU	ıyı	GIII	75	PILE	
agg	gag	aat	gtg	cta	cga	aac	cta	gcg	gat	aag	gcc	ttt	gac	cgg	CCC	481
Arg	Glu	Asn	Val	Leu	Arg	Asn	Leu	Ala	Asp	Lys	Ala	Phe	Asp	Arg	Pro	
			80					85					90			529
atc	tgc	gag	gcc	ctc	ctg	gac	cag	agg	ttc	ttc	aat	ggc	att	990	aac	323
Ile	Cys	Glu	Āla	Leu	Leu	Asp		Arg	Phe	Phe	Asn	GIY	TTE	GIY	ASII	
		95					100					105				E 77
tat	ctg	cgg	gca	gag	atc	ctg	tac	cgg	ctg	aag	atc	ccc	ccc	-:	gag	577
Tyr	Leu 110	Arg	Ala	Glu	Ile	Leu 115	Tyr	Arg	Leu	Lys	Ile 120	Pro	Pro	Pne	GIU	
			tcg	at a	ata		acc	cta	cad	cad		agg	cca	agc	cca	625
aag	33-	ege	Ser	37-3	tou	949	מות	Len	Gla	Gln	His	Ara	Pro	Ser	Pro	
	Ala	Arg	ser	vai	130	Gru	Ala	nea	GIII	135	*****	*** 9			140	
125								-~-			at a	CaG	aat	cca		673
gag	ctg	acc	ctg	agc	cag	aag	ata	agg	The	Tara	Tou	Cla	Acn	Dro	Agn	
GIu	Leu	Thr	Leu		GIN	гÀа	TIE	Arg		цув	neu	Gin	No.	155	rob	
				145					150			~+ à	a.a.		aaa	721
ctg	ctg	gag	cta	tgt	cac	tca	grg	ccc	aag	gaa	grg	gre	cag	Tay	999	, 2 1
Leu	Leu	Glu	Leu	Cys	His	Ser	Val		гÀг	GIU	Val	Vai	170	пеα	Giy	
			160					165								769
ggc	aga	ggc	tac	ggg	tca	gag	agc	<u>aāa</u>	gag	gag	gac	ttt	gct	gcc	27.	103
Gly	Arg	Gly	Tyr	Gly	Ser	Glu	Ser	Gly	Glu	Glu	Asp		Ala	АТА	Pne	
		175					180					185				017
cga	gcc	tgg	ctg	cgc	tgc	tat	ggc	atg	cca	ggc	atg	agc	tcc	ctg	cag	817
Arg	Ala	Trp	Leu	Arg	Cys	Tyr	Gly	Met	Pro	Gly	Met	Ser	ser	Leu	Gin	
	190					195					200					0.55
gac	cgg	cat	ggc	cgt	acc	atc	tgg	ttc	cag	999	gat	cct	gga	ccg	ttg	865
Asp	Arg	His	Gly	Arg	Thr	Ile	Trp	Phe	Gln	Gly	Asp	Pro	Gly	Pro	Leu	
205					210					215					220	
gca	ccc	aaa	999	cgc	aag	tcc	cgc	aaa	aag	aaa	tcc	aag	gcc	aca	cag	913
Āla	Pro	Lys	Gly	Arg	Lys	Ser	Arg	Lys	Lys	Lys	Ser	Lys	Ala	Thr	Gln	
				225					230					235		
ctq	aqt	cct	gag	gac	aga	gtg	gag	gac	gct	ttg	cct	ccg	agc	aag	gcc	961
Leu	Ser	Pro	Glu	Asp	Arq	Val	Glu	Asp	Ala	Leu	Pro	Pro	Ser	Lys	Ala	
			240	-	_			245					250			
cct	tcc	agg	aca	cqa	agg	gca	aag	aga	gac	ctt	cct	aag	agg	act	gca	1009
Pro	Ser	Arg	Thr	Arq	Arq	Ala	Lys	Arg	Asp	Leu	Pro	Lys	Arg	Thr	Ala	
		255			_		260		_			265				
acc	caq		cct	qaq	ggg	acc	agc	ctc	cag	cag	gac	cca	gaa	gct	CCC	1057
Thr	Gln	Arq	Pro	Glu	Gly	Thr	Ser	Leu	Gln	Gln	Asp	Pro	Glu	Ala	Pra	
	270				•	275					280					
aca	ata	ccc	aag	aaq	aaa	agg	agg	aag	ggg	cga	cag	gca	gcc	tct	ggc	1105
Thr	Val	Pro	Lvs	Lvs	Gly	Arg	Arg	Lys	Gly	Arg	Gln	Ala	Ala	Ser	Gly	
285			-1-	-3-	290		_	•	•	295					300	
cac	tac	aga	ccc	caa			aaq	act	gac	ato	сса	tcc	ttg	gaa	cca	1153
Hie	Cve	Aro	Pro	Ara	Lvs	Val	Lvs	Ala	Asp	Ile	Pro	Ser	Leu	Glu	Pro	
	-, -	3		305			-2-		310					315		
gag	aaa	acc	tca			tag	cago	aga			ac t	tgca	ctca	C		1201
			Ser				~~33	-33			٠ - در	ے کی کے ح				
GIU	. Сту	1111	320		361			**								
~~-	++	t a #				ace +	ctoo	ום מת	tete	aa++		aaaa	aca	gaca	atatct	1261
727	act c	cat	2000	acca	te c	gac+		צב כי לה הי	gcac	aact	. ctc	aton	ttt	taat	tgtacc	1321
yaa	#4C	caa	acag	9000	22 -		+~+~	ia en	3240	, auc	. 200	33 tt:	ata	aact	gataca	1381
					aa 9	, Luca	وعودي	a aa	واعمد	Junge					J	1398
ככנ	yaaa	aaa	aaaa	aaa												,

```
<210> 72
<211> 821
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 42..611
<221> sig_peptide
<222> 42..287
<223> Von Heijne matrix
     score 4.4
     seq NLPHLQVVGLTWG/HI
<221> polyA_signal
<222> 787..792
<221> polyA_site
<222> 808..821
<400> 72
cogttgccag ttctgcgcgt gtcctgcgtc tccagtatgg a atg tat gtt tgg ccc
                                                                       56
                                              Met Tyr Val Trp Pro
                                                       -80
tgt gct gtg gtc ctg gcc cag tac ctt tgg ttt cac aga aga tct ctg
Cys Ala Val Val Leu Ala Gln Tyr Leu Trp Phe His Arg Arg Ser Leu
        -75
                            -70
                                                -65
cca ggc aag gcc atc tta gag att gga gca gga gtg agc ctt cca gga
                                                                      152
Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly Val Ser Leu Pro Gly
                                            -50
                        - 55
att ttg act gcc aaa tgt ggt gca gaa gta ata ctg tca gac agc tca
Ile Leu Thr Ala Lys Cys Gly Ala Glu Val Ile Leu Ser Asp Ser Ser
-45
                                        -35
gaa ctg cct cac tgt ctg gaa gtc tgt cgg caa agc tgc caa atg aat
                                                                      248
Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln Ser Cys Gln Met Asn
                                    -20
                -25
                                                                      296
aac ctg cca cat ctg cag gtg gta gga cta aca tgg ggt cat ata tct
Asn Leu Pro His Leu Gln Val Val Gly Leu Thr Trp Gly His Ile Ser
                                -5
tgg gat ctt ctg gct cta cca cca caa gat att atc ctt gca tct gat
                                                                      344
Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile Ile Leu Ala Ser Asp
gtg ttc ttt gaa cca gaa gat ttt gaa gac att ttg gct aca ata tat
                                                                      392
Val Phe Phe Glu Pro Glu Asp Phe Glu Asp Ile Leu Ala Thr Ile Tyr
                                        30
                                                            35
                                                                      440
ttt ttg atg cac aag aat ccc aag gtc caa ttg tgg tct act tat caa
Phe Leu Met His Lys Asn Pro Lys Val Gln Leu Trp Ser Thr Tyr Gln
                40
                                    45
gtt agg agt gct gac tgg tca ctt gaa gct tta ctc tac aaa tgg gat
                                                                      488
Val Arg Ser Ala Asp Trp Ser Leu Glu Ala Leu Leu Tyr Lys Trp Asp
           55
                                60
                                                    65
                                                                      536
atg aaa tgt gtc cac att cct ctt gag tct ttt gat gca gac aaa gaa
Met Lys Cys Val His Ile Pro Leu Glu Ser Phe Asp Ala Asp Lys Glu
                            75
                                                80
                                                                      584
gat ata gca gaa tot acc ott cca gga aga cat aca gtt gaa atg otg
Asp Ile Ala Glu Ser Thr Leu Pro Gly Arg His Thr Val Glu Met Leu
                        90
gtc att tcc ttt gca aag gac agt ctc tgaattatac ctacaacctg
Val Ile Ser Phe Ala Lys Asp Ser Leu
```

•								
100 ttctgggaca cagcttgaga cacctagaca aaaaaaaaaa	gtatcaata	g tctgaaga	atg gtca	agtctg t	ctgcctta	g accers	gacyc	691 751 811 821
<210> 73								
<211> 916								
<212> DNA <213> Homo	sapiens	***						
<220> <221> CDS								
<222> 629	16							
<221> sig_p <222> 627								
<223> Von H	Reijne mat	rix						
	LVTPAALRPL	VLG/GN						
<221> poly# <222> 904								
<400> 73 cctgaatgac	ttgaatgtt	t ccccqcc	tga gcta	aacagtc	catgtggg	g attca	gctct	60
g atg gga t Met Gly (gt gtt tt Cys Val Ph	c cag age	aca gaa	a gac aa	a cgt ata	e Phe Ly	g ala	109
gac tgg act Asp Trp Th	t ctg tca r Leu Ser	Pro Gly G -210	lu His 1	Ala Lys	Asp Glu : -205	ryr var	Leu	157
tac tat tac Tyr Tyr Ty:	tcc aat r Ser Asn	Leu Ser V	tg cct a al Pro :	Ile Gly	Arg Phe	cag aac Gln Asn	cgc Arg -185	205
-200 gta cac tte	a ata aga	-195	ta toc a	-190 aat gat		ctc ctg		253
Val His Le	u Met Gly -180	Asp Asn L	eu Cys i	Asn Asp -175	Gly Ser	Leu Leu -170	neu)	
caa gat gt	g caa gag	gct gac c	ag gga a	acc tat Thr Tvr	atc tgt Ile Cvs	gaa atc Glu Ile	cgc Arg	301
	-165		-160			-155		349
ctc aaa gg Leu Lys Gl	y Glu Ser	Gln Val F	tc aag Phe Lys : -145	aag gcg Lys Ala	Val Val -140	Leu His	Val	3.2
-1 ctt cca ga	a gag ccc	aaa qaq q	ctc atg	gtc cat	gtg ggt	gga ttg		397
Leu Pro Gl -135	u Glu Pro	Lys Glu I -130	Leu Met	Val His	-125	GIA ren	116	445
cag atg gg Gln Met Gl	a tgt gtt y Cys Val	ttc cag a Phe Gln S -115	agc aca Ser Thr	gaa gtg Glu Val -110	Lys His	gtg acc Val Thr	aag Lys -105	445
-120 gta gaa tg	g ata ttt	tca qqa q	egg ege	gca aag	gag gag	att gta	ttt	493
Val Glu Tr	p Ile Phe -10	Ser Gly 1 0	Arg Arg	Ala Lys -95	Glu Glu	-90	Pne	C 4.3
cgt tac ta Arg Tyr Ty	r His Lys	ctc agg a	Met Ser	gcg gag Ala Glu	tac tcc Tyr Ser	cag agc Gln Ser -75	tgg Trp	541
ggc cac tt	-85 c cag aat	cgt gtg a	-80 aac ctg	gtg ggg	gac att	ttc cgc	aat	589
Gly His Ph	ne Gln Asn 70	.Arg Val.	Asn Leu -65	Val Gly	-60	Phe Arg	ASII	627
gac ggt to	c atc atg	ctt caa	gga gtg	agg gag	tca gat	gga gga	aac	637

Asp																
	Gly	Ser	Ile	Met	Leu	Gln -50	Gly	Val	Arg	Glu	Ser	Asp	Gly	Gly	Asn	
tac		tac	aqt	atc	cac		999	aac	ctg	gtg	ttc	aag	aaa	acc	att	685
Tvr	Thr	Cys	Ser	Ile	His	Leu	Gly	Asn	Leu	Val	Phe	Lys	Lys	Thr	Ile	
-40		- 2 -			-35		•			-30		•			-25	
	cta	cat	qtc	agc	ccq	gaa	gag	cct	cga	aca	ctg	gtg	acc	ccg	gca	733
Val	Leu	His	Val	Ser	Pro	Glu	Glu	Pro	Arq	Thr	Leu	Val	Thr	Pro	Ala	
				-20					-15					-10		
acc	cta	agg	cat		atc	tta	ggt	aat	aat	caq	tta	ata	atc	att	gtg	781
Ala	Leu	Ara	Pro	Leu	Val	Leu	Gly	Glv	Asn	Gln	Leu	Val	Ile	Ile	Val	
n.z.u		••• 5	-5					1				5				
ada	att	atc		gcc	aca	atc	ctg		ctc	cct	atc	ctq	ata	ttq	atc	829
olv.	Tle	Val	Cvs	Ala	Thr	Tle	Leu	Leu	Leu	Pro	Val	Leu	Ile	Leu	Ile	
- 1	10		-1-		-,	15					20					
ta		aaq	acc	tat	gga		aag	agt	tca	ata		tct	aca	qtc	ttq	877
7=1	Lve	Live	Thr	CVS	Glv	Asn	Lys	Ser	Ser	Val	Asn	Ser	Thr	Val	Leu	
5	Lys	<i>D</i>		Cy D	30		_,_			35					40	
	220	aac	acq.	aad		act	aat	cca	aaa		aaa	aaa				916
, - y	Lve	Aen	アアン	Live	Live	Thr	Asn	Pro	Lvs	Lvs	Lvs	Lvs				
41	my 5	veii	****	15 45	-y 5				50	_, ~	-,-	-,-				
<21	0 > 74	<u>1</u>														
	1> 1:															
	2 > D1															
		omo s	sapie	ens					,							
			_													
<22	0 >															
<22	1> CI	os														
<22	2 > 62	252	20													
				_												
<22	1> po	olyA_	_sigr	nai												
		olyA_ 124	-													
			-													
<22	2 > 1:		1129	€												
<22 <22	2 > 1: 1 > po	124.	.1129 _site	€												,
<22 <22	2 > 1: 1 > po	124. olyA_	.1129 _site	€												,
<22: <22: <22: <40:	2 > 1: 1 > po 2 > 1: 0 > 7	124. olyA_ 141.	.1129 _site	e 3												,
<22: <22: <22: <40: cct;	2 > 1: 1 > po 2 > 1: 0 > 74 gaats	124. olyA 141. 4 gac t	.1129 _site .1153	e 3 atgt:	tt c	ccg	cctga	a gct	taaca	agtc	cato	gtgg	gtg a	atto	agctct	60
<22: <22: <22: <40: cct: g a:	2 > 1: 1 > po 2 > 1: 0 > 74 gaats	olyA 141. 4 gac t	site site 1153	e 3 atgt:	tc ca	ag ag	gc a	ca gi	ta ga	ac aa	aa to	gt at	a t	c a	ag ata	60 109
<22: <22: <22: <40: cct: g a:	2 > 1: 1 > po 2 > 1: 0 > 74 gaats	olyA 141. 4 gac t	site site 1153	e 3 atgt:	tc ca	ag ag	gc a	ca gi	ta ga	ac aa	aa to	gt at	a t	c a	agctct ag ata ys Ile	
<22: <22: <22: <40: cct: g a: M:	1> po 1> po 2> 1: 0> 74 gaato tg go et G	olyA olyA 141. 4 gac t ga t gly C	_site _site .115: ctgaa gt gt	atgtt tt tt al Pl	tc ca ne Gi	ag ag ln Se	gc ad er Tl	ca gi nr Va	ta ga al Aa 10	ac aa sp Ly O	aa to /s Cy	gt at /s I	ta ti le Pl	c and a line Line Line Line Line Line Line Line L	ag ata ys Ile 5	109
<22: <22: <22: <40: cct: g a: M:	1> po 2> 1: 0> 74 gaate tg ge et G	olyA 141. 4 gac t ga to	_site _site .115: ctgaa gt gt ys Va	atgt: tt tt al Pl 5	tc ca ne Gi cca	ag ag ln Se gga	gc acer Tl	ca gi nr Va cac	ta ga al Aa 10 gcc	ac aa sp Ly O aag	aa to ys Cy gac	gt at /s I: gaa	ta ti le Pl tat	tc and the Lyngh 1!	ag ata ys Ile 5 cta	
<22: <22: <22: <40: cct: g a M	1> po 2> 1: 0> 74 gaate tg ge et G	olyA 141. 4 gac t ga to	_site _site .115: ctgaa gt gt ys Va	atgt: tt tt al Pl 5	tc ca ne Gi cca	ag ag ln Se gga	gc ad er Tl	ca gi nr Va cac	ta ga al Aa 10 gcc	ac aa sp Ly O aag	aa to ys Cy gac	gt at /s I: gaa	ta ti le Pl tat	tc and the Lyngh 1!	ag ata ys Ile 5 cta	109
<22: <22: <22: <40: cct: g a M	1> po 2> 1: 0> 74 gaate tg ge et G	olyA 141. 4 gac t ga to	_site _site .115: ctgaa gt gt ys Va	atgt: tt tt al Pl 5	tc ca ne Gi cca	ag ag ln Se gga	gc acer Tl	ca gi nr Va cac	ta ga al Aa 10 gcc	ac aa sp Ly O aag	aa to ys Cy gac	gt at /s I: gaa	ta ti le Pl tat	tc and the Lyngh 1!	ag ata ys Ile 5 cta	109
<22: <22: <40: cct; g a M 1 gac Asp	1> po 2> 1: 0> 7. gaat; tg gg et G tgg	olyA olyA 141. 4 gac t ga t ga t ga t Thr	site site 1153 ttgaa gt gt ys Va ctg Leu 20	atgti tt ti al Pl 5 tca Ser	cca Pro	ag ag ln Se gga Gly	gc ac er Tl gag Glu	ca go nr Va cac His 25	ta ga al As al a gcc Ala	ac aa sp Ly 0 aag Lys	gac Asp	gt at /s II gaa Glu	ta ti le Pl tat Tyr 30	e a ne Ly 1! gtg Val	ag ata ys Ile 5 cta Leu	109
<22: <22: <40: cct: g a: 1: gac Asp	2 > 1: 1 > po 2 > 1: 0 > 7. gaato tg go et G: tgg Trp	olyA olyA l41. 4 gac t ga t gly Cy act Thr	site site 1153 tgaa gt gt ys Va ctg Leu 20 tcc	atgti tt ti al Pl 5 tca Ser	tc ca ne G cca Pro	ag ag ln Se gga Gly agt	gc acer Ti gag Glu gtg	ca go cac His 25 cct	ta ga al As gcc Ala att	ac aasp Lys aag Lys	gac Asp	gt at /s I: gaa Glu ttc	tat tat Tyr 30 cag	c and	ag ata ys Ile 5 cta Leu cgc	109
<22: <22: <40: cct: g a: 1: gac Asp	2 > 1: 1 > po 2 > 1: 0 > 7. gaato tg go et G: tgg Trp	olyA olyA olyA oly C ga to ga to ga to tac Thr tac	site site 1153 tgaa gt gt ys Va ctg Leu 20 tcc	atgti tt ti al Pl 5 tca Ser	tc ca ne G cca Pro	ag ag ln Se gga Gly agt	gc ac er Tl gag Glu	ca go cac His 25 cct	ta ga al As gcc Ala att	ac aasp Lys aag Lys	gac Asp	gt at /s I: gaa Glu ttc	tat tat Tyr 30 cag	c and	ag ata ys Ile 5 cta Leu cgc	109
<222 <222 <400 cct. g a M. 1 gac Asp	2 > 1: 1 > po 2 > 1: 0 > 7. gaatg tg gg et G: tgg Trp tat Tyr	olyA olyA olyA oly C ga to ga to ga to tac Thr tac Tyr 35	_site _site .1153 ctgaa gt gt ys Va ctg Leu 20 tcc Ser	atgt: tt t: al P! tca Ser aat Asn	cca Pro	gga Gly agt Ser	gc acer Tl gag Glu gtg Val 40	cac His 25 cct Pro	ta ga al As gcc Ala att Ile	ac aa sp Ly aag Lys ggg Gly	gac Asp cgc Arg	gaa Glu ttc Phe 45	tat Tyr 30 cag	gtg Val	ag ata ys Ile tota Leu cgc Arg	109
<222 <222 <400 ccct g a M 1 gac Asp tac Tyr	2 > 1: 1 > po 2 > 1: 0 > 7. gaat; tg gg et G: tgg Trp tat Tyr	olyA l41. 4 gac t ga t gly C act Thr tac Tyr 35 ttg	_site _site _115: ctgaa gt gt ys Va ctg Leu 20 tcc Ser atg	atgti atgti at ti al Pi tca Ser aat Asn	cca Pro ctc Leu gac	ag ag ln So gga Gly agt Ser	gc acer Tl gag Glu gtg Val 40 tta	ca gi cac His 25 cct Pro	ta gal As al As gcc Ala att Ile aat	ac as sp Ly aag Lys ggg Gly	gac Asp cgc Arg	gaa Glu ttc Phe 45	tat Tyr 30 cag Gln	tc and left	ag ata ys Ile tota Leu cgc Arg	109 157 205
<222 <222 <400 cct gam h gac Asp tac Tyr	2 > 1: 1 > po 2 > 1: 0 > 74 gaats tg gs et G: tgg Trp tat Tyr cac His	olyA l41. 4 gac t ga t gly C act Thr tac Tyr 35 ttg	_site _site _115: ctgaa gt gt ys Va ctg Leu 20 tcc Ser atg	atgti atgti at ti al Pi tca Ser aat Asn	cca Pro ctc Leu gac	gga Gly agt Ser	gc acer Tl gag Glu gtg Val 40	ca gi cac His 25 cct Pro	ta gal As al As gcc Ala att Ile aat	ac as sp Ly aag Lys ggg Gly	gac Asp cgc Arg ggc Gly	gaa Glu ttc Phe 45	tat Tyr 30 cag Gln	tc and left	ag ata ys Ile tota Leu cgc Arg	109 157 205
<22 <22 <40 cct M 1 gac Asp tac Tyr gtal	2 > 1: 1 > po 2 > 1: 0 > 7. gaatg tg gg et G: Trp tat Tyr cac His	olyA 141. 4 gac t ga t gly C act Thr tac Tyr 35 ttg Leu	site sitgaa gt gt ys Va ctg Leu 20 tcc Ser atg	atgti at ti al Pi tca Ser aat Asn 999	cca Pro ctc Leu gac Asp	gga Gly agt Ser atc Ile	gc acer Tl gag Glu gtg Val 40 tta Leu	cac His 25 cct Pro tgc	ta ga al As gcc Ala att Ile aat Asn	ac as sp Ly aag Lys ggg Gly gat Asp	gac Asp cgc Arg ggc Gly 60	gaa Glu ttc Phe 45 tct Ser	tat Tyr 30 cag Gln ctc Leu	gtg Val aac Asn ctg Leu	ag ata ys Ile tota Leu cgc Arg ctc Leu	109 157 205
<222 <222 <40 cct. g a M 1 gacAsp tac Tyr gtal Val	2 > 1: 1 > po 2 > 1: 0 > 7 gaatg et g Trp tat Tyr cac His 50 gat	olyA 141. 4 gac t ga t ga t gly C act Thr tac Tyr 35 ttg Leu	site site site site site site site site	atgtitied pi	cca Pro ctc Leu gac Asp	gga Gly agt ser atc Ile 55 gac	gc acer Tl gag Glu gtg Val 40 tta Leu cag	cac His 25 cct Pro tgc Cys	ta ga al As gcc Ala att Ile aat Asn	ac as sp Ly aag Lys ggg Gly gat Asp	gac Asp cgc Arg ggc Gly 60 atc	gaa Glu ttc Phe 45 tct Ser	tat Tyr 30 cag Gln ctc Leu	gtg Val aac Asn ctg Leu	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc	109 157 205 253
<22 <22 <40 cct ga M l gac Asp tac Tyr gtal Caa Gln	2 > 1: 1 > po 2 > 1: 0 > 7 gaatg et g Trp tat Tyr cac His 50 gat	olyA 141. 4 gac t ga t ga t gly C act Thr tac Tyr 35 ttg Leu	site site site site site site site site	atgtitied pi	cca Pro ctc Leu gac Asp	gga Gly agt ser atc Ile 55 gac	gc acer Tl gag Glu gtg Val 40 tta Leu	cac His 25 cct Pro tgc Cys	ta ga al As gcc Ala att Ile aat Asn	ac aasp LyD aag Lys ggg Gly gat Asp tat Tyr	gac Asp cgc Arg ggc Gly 60 atc	gaa Glu ttc Phe 45 tct Ser	tat Tyr 30 cag Gln ctc Leu	gtg Val aac Asn ctg Leu	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg	109 157 205 253
<222 <222 <400 cctc g a M 1 gacpAsp tacrTyr gtal Val caa GGln 65	2 > 1: 1 > po 2 > 1: 0 > 7: gaatg tg gg tt gg Trp tat Tyr cac His 50 gat Asp	olyA l41. 4 gac t ga t gly C act Thr tac Tyr 35 ttg Leu gtg Val	site site site site site site site site	atgti at ti al Pi tca Ser aat Asn ggg Gly gag	cca Pro ctc Leu gac Asp gct Ala	gga gga Gly agt ser atc Ile 55 gac Asp	gc acer Tl gag Glu gtg Val 40 tta Leu cag Gln	ca gt cac His 25 cct Pro tgc Cys	ta gal As lo gcc Ala att Ile aat Asn acc Thr	ac aasp LyD aag Lys ggg Gly gat Asp tat Tyr 75	gac Asp cgc Arg ggc Gly 60 atc	gaa Glu ttc Phe 45 tct Ser tgt Cys	tat Tyr 30 cag Gln ctc Leu gaa Glu	gtg Val aac Asn ctg Leu	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg scale cgc Arg	109 157 205 253
<222 <222 <400 cctc g a M 1 gacpAsp tacrTyr gtal caa GGln 65 cctc	2 > 1: 1 > po 2 > 1: 0 > 7. gaa gg et t gg Trp tatr Cac His 50 at Asp	olyA l41. 4 gac tga tga ly Cy act Thr tac Tyr 35 ttg Leu gtg Val	site site site site site site site site	atgtitied plants of the ser and ser and ser and ser ggg gly gag glu agc	cca Pro ctc Leu gac Asp gct Ala 70 cag	gga gga Gly agt ser atc Ile 55 gac Asp	gc acer Tl gag Glu gtg Val 40 tta Leu cag Gln ttc	ca gthat Value of the Value of	ta gal As al As gcc Ala att Ile aat Asn acc Thr	ac aasp Lyo aag Lys ggg Gly gat Asp tat Tyr 75 gcg	gac Asp cgc Arg ggc Gly 60 atc Ile gtg	gaa Glu ttc Phe 45 tct Ser tgt Cys	tat Tyr 30 cag Gln ctc Leu gaa Glu ctg	gtg Val aac Asn ctg Leu atc	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg gtg	109 157 205 253
<222 <222 <400 cctc g a M 1 gacpAsp tacrTyr gtal caa GGln 65 cctc	2 > 1: 1 > po 2 > 1: 0 > 7. gaa gg et t gg Trp tatr Cac His 50 at Asp	olyA l41. 4 gac tga tga ly Cy act Thr tac Tyr 35 ttg Leu gtg Val	site site site site site site site site	atgtt at tt al Pl 5 tca Ser aat Asn ggggly gag Glu agc	cca Pro ctc Leu gac Asp gct Ala 70 cag	gga gga Gly agt ser atc Ile 55 gac Asp	gc acer Tl gag Glu gtg Val 40 tta Leu cag Gln	ca gthat Value of the Value of	ta gal As logcc Ala att Ile aat Asn acc Thr aag Lys	ac aasp Lyo aag Lys ggg Gly gat Asp tat Tyr 75 gcg	gac Asp cgc Arg ggc Gly 60 atc Ile gtg	gaa Glu ttc Phe 45 tct Ser tgt Cys	tat Tyr 30 cag Gln ctc Leu gaa Glu ctg	gtg Val aac Asn ctg Leu atc Ile cat	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg gtg	109 157 205 253
<222 <222 <400 cctc g a M 1 gacpAsp tacrTyr gtal Caa GGln 65 ctc Leu	2 > 1: 1 > po 2 > 1: 0 > 7: gaa gg et t gg Trp tatr Cac His 50 at Asp aaa Lys	olyA l41. 4 gac tga tga ly Cy act Thr tac Tyr 35 ttg Leu gtg Val	site site site site site site site site	atgtities to the ser as a ser	cca Pro ctc Leu gac Asp gct Ala 70 cag Gln	gga Gly agt Ser atc Ile 55 gac Asp Val	gc acer Tl gag Glu gtg Val 40 tta Leu cag Gln ttc Phe	cae His 25 cct Pro Cys gga Gly aag Lys	ad Asi gcc Ala att Ile aat Asn acc Thr aag Lys	ac aasp Lyo aag Lys ggg Gly gat Asp tat Tyr 75 gcg Ala	gac Asp cgc Arg ggc Gly 60 atc Ile gtg Val	gt at ys I: gaa Glu ttc Phe 45 tct Ser tgt Cys gta Val	tat Tyr 30 cag Gln ctc Leu gaa Glu ctg	gtg Val aac Asn ctg Leu atc Ile cat His	ag ata ys lle tota Leu cgc Arg ctc Leu cgc Arg sqc Arg val	109 157 205 253 301 349
<222 <222 <400 cct gas M 1 cgas tryr tal aan 665 cct ctt	2 > 1: 1 > po 2 > 1: 0 > at: 0 > at: 0 = t t t t t t t t t t t t t t t t t t	olyA l41. 4 gac tga ga tga ly Cy act Thr tac Tyr 35 ttg Val ggg Val	site stgaa gt gt ys Va ctg Leu 20 tcc Ser atg Met caa Gln gag	atgtital plant of the ser at Asn ggly gag Glu ager 85 cc	cca Pro ctc Leu gac Asp gct Ala 70 cag Gln	ag aglan Sollan	gc acer Ti gag Glu gtg Val 40 tta Leu cag Gln ttc Phe ctc	cac His 25 cct Pro tgc Cys gga Gly aag Lys	ta gal As att Ile aatt Asn acc Thr aag Lys	ac aasp Lyo aag Lys ggg Gly gat Asp tat Tyr 75 gcg Ala cat	gac Asp cgc Arg ggc Gly 60 atc Ile gtg Val gtg	gt at ys II gaa Glu ttc Phe 45 tct Ser tgt Cys gtal ggt	tat Tyr 30 cag Gln ctc Leu gaa Glu ctg Leu gga	gtg Val aac Asn ctg Leu atc Lis 95 ttg	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg stg Val att	109 157 205 253
<222 <222 <400 cctc g a M l cgAsp tTyr tal acG65 ccLeu ctt	2 > 1: 1 > po 2 > 1: 0 > at: 0 > at: 0 = t t t t t t t t t t t t t t t t t t	olyA l41. 4 gac tga ga tga ly Cy act Thr tac Tyr 35 ttg Val ggg Val	site strategy very solution of the s	atgtital plant of the ser at Asn ggly gag Glu ager 85 cc	cca Pro ctc Leu gac Asp gct Ala 70 cag Gln	ag aglan Sollan	gc acer Tl gag Glu gtg Val 40 tta Leu cag Gln ttc Phe	cac gthis vertical ve	ta gal As att Ile aatt Asn acc Thr aag Lys	ac aasp Lyo aag Lys ggg Gly gat Asp tat Tyr 75 gcg Ala cat	gac Asp cgc Arg ggc Gly 60 atc Ile gtg Val gtg	gt at ys II gaa Glu ttc Phe 45 tct Ser tgt Cys gtal ggt	tat Tyr 30 cag Gln ctc Leu gaa Glu ctg Leu gga Gly	gtg Val aac Asn ctg Leu atc Lis 95 ttg	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg stg Val att	109 157 205 253 301
<222 <222 <400 cctc g a M 1 gasp tayr tayr tal cGln 655 ccLeu cttu	2 > 1: 1 > po 2 > 1: 0 > 7: gaa gg tet gg Trp tarr cas Sgat Asp aaa Lys ca	olyA l41. 4 gac tga ga tga ly Cy act Thr tac Tyr 35 ttg Val ggg Gly gag Glu	site site site site site site site site	atgtital plant of the ser at Asn gggy gag Glu agc ser ecc	cca Pro ctc Leu gac Asp gct Ala 70 cag Gln aaa Lys	gga Gly agt Ser atc Ile 55 gac Asp Val gag Glu	gc acer Ti gag Glu gtg Val 40 tta Leu cag Gln ttc Phe ctc Leu	cac His 25 cct Pro Cys gga Gly aag Lys atg	ad Asi gcc Ala att Ile aat Asn acc Thr aag Lys 90 gtc Val	ac aasp Lyo aag Lys ggg Gly gat Asp tat Tyr 75 gcg Ala cat His	gac Asp cgc Arg ggc Gly 60 atc Ile gtg Val	gt at ys II gaa Glu ttc Phe 45 tct Ser tgt Cys gta Val ggt Gly	tat Tyr 30 cag Gln ctc Leu gaa Glu ctg Leu gga Gly 110	gtg Val aac Asn ctg Leu atc Lis 55 ttg Leu	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg stg Val att Ile	109 157 205 253 301 349
<222 <222 <400 cctc g a Mi 1 gac Asp tac Tyr gta Val caa Gln 65 ctc Leu ctt Leu	2 > 1: 1 > po 2 > 1: 0 > 7: gaa gg tet gg Trp tarr cas Sgat Asp aaa Lys ca	olyA l41. 4 gac tga ga tga ly Cy act Thr tac Tyr 35 ttg Val ggg Gly gag Glu	site site site site site site site site	atgtital plant of the ser at Asn gggy gag Glu agc ser ecc	cca Pro ctc Leu gac Asp gct Ala 70 cag Gln aaa Lys	gga Gly agt Ser atc Ile 55 gac Asp Val gag Glu	gc acer Ti gag Glu gtg Val 40 tta Leu cag Gln ttc Phe ctc	cac His 25 cct Pro Cys gga Gly aag Lys atg	ad Asi gcc Ala att Ile aat Asn acc Thr aag Lys 90 gtc Val	ac aasp Lyo aag Lys ggg Gly gat Asp tat Tyr 75 gcg Ala cat His	gac Asp cgc Arg ggc Gly 60 atc Ile gtg Val	gt at ys II gaa Glu ttc Phe 45 tct Ser tgt Cys gta Val ggt Gly	tat Tyr 30 cag Gln ctc Leu gaa Glu ctg Leu gga Gly 110	gtg Val aac Asn ctg Leu atc Lis 55 ttg Leu	ag ata ys Ile tota Leu cgc Arg ctc Leu cgc Arg stg Val att Ile	109 157 205 253 301

Gln Met Gly Cys Val Phe Gln Ser Thr Glu Val Lys His Val Thr Lys 115 120 125	400 %
and the state of the transparence of the state of the sta	493 .
Val Glu Trp Ile Phe Ser Gly Arg Arg Ala Lys Val Thr Arg Arg Lys	
cat cac tgt gtt aga gaa ggc tct ggc tgatggtatc aggacaaagg	540
His His Cys Val Arg Glu Gly Ser Gly	
150	600
tagaatcagg cacatgagga ggtgttgcaa gagcctgggc tttggtgctt atcagaactg gaccttctcc tagcaatttc agctttctgg tgggaaaggt aactccaatg aagaacaaga	660
	720 780
	840
water atomics atomics of managed at the telegraphic constraints	900
gctagacatt aaaatgatta cacttttatt cttactgggg ttagttctgt gagttttcat ctgtgcccca ttgccccatt tatgtgatgg agggaatttt catgggtact tcacgtgttg	960
	1020
	1080 1140
gggtcaacat gtgttgtggg gatatcccaa gtaactigtt actaataaaa gtdaggog	1153
aaaaaaaaa aaa	
\cdot	
<210> 75	
<211> 1517	
<212> DNA <213> Homo sapiens	
<220>	
<221> CDS <222> 21167	
<2225 21167	
<400> 75	53
ctctgaaatg cttgtctttt atg ctg gna ggt gac cat agg gct ctg ctt tta	
Mak Tay Vas CIV Ach His Ard Ald Ded Ded Ded	
Met Leu Xaa Gly Asp His Arg Ala Leu Leu Leu 1 5 10	
1 5 10 20 20 20 20 20 20 20 20 20 20 20 20 20	101
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro	101
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 20 25	101 149
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25	
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40	149
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 ggg aga tta gtg gtg atg gag aga ggt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct	
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys	149
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45	149 197 257
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 caactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt	149 197 257 317
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcagagttgt acatatgttt	149 197 257 317 377
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcagagttgt acatatgttt cacagtacag gatctgtaca taaaagtttc tttcctaaac cattcaccaa gagccaatat ctaggcattt tettggtagc acaaattttc ttattgctta gaaaattgtc ctccttgtta	149 197 257 317
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcagagttgt acatatgttt cacagtacag gatctgtaca taaaagttc tttctctaaac cattcaccaa gagccaatat ctaggcattt tcttggtagc acaaattttc ttattgctta gaaaattgtc ctccttgtta tttctgtttg taagacttaa gtgagtagg tctttaagga aagcaacgct cctctgaaat tttctgtttg taagacttaa gtgagtagg tctttaagga aagcaacgct cctctgaaat	149 197 257 317 377 437 497 557
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcagagttgt acatatgttt cacagtacag gatctgtaca taaaagttc tttctaaac cattcacaa gagccaatat ctaggcattt tcttggtagc acaaattttc ttattgctta gaaaattgtc ctccttgtta ttctgtttg taagacttaa gtgagttagg tctttaagga aagcaacgct cctctgaaat gcttgtcttt tatgctggga ggtgaccata gggctctgct tttaaagaa ttggctgcttc aagggcaga gtcaacaag ggacttcttc cagggagatt agtggtgatg gagaggagag	149 197 257 317 377 437 497 557 617
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcagagttgt acatatgttt cacagtacag gatctgtaca taaaagttc tttctaaac cattcacaa gagccaatat ctaggcattt tcttggtagc acaaatttc ttattgctta gaaaattgtc ctccttgtta ttctgtttg taagacttaa gtgagtagg tctttaagga aagcaacgct cctctgaaat gcttgtctt tatgctgga ggtgaccata gggctctgct tttaaagaa ttggctgctc aaaggccaga gtcacaggaa ggacttcttc cagggagatt agtggtgatg gagaggagag	149 197 257 317 377 437 497 557 617 677
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcagagttgt acatatgttt cacagtacag gatctgtaca taaaagttc tttcctaaac cattcaccaa gagccaatat ctaggcattt tcttggtagc acaaatttc ttattgctta gaaaattgtc ctccttgtta tttctgtttg taagacttaa gtgagttagg tctttaagga aagcaacgct cctctgaaat gcttgtctt tatgctgga ggtgaccata gggctctgct tttaaagaa tggctgctct aaaggccaga gtcacaggaa ggtgaccata gggctctgct tctaaagata tggctgctc aagggcaga ctcatgtcct tcttgtccac ggttttgttg agttttcact cttctaatgc aagggcaga ctctatgcca cacttaggat qtgatcactt tccaggtggcc aggaaggagag	149 197 257 317 377 437 497 557 617
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgttt tggctcagtt catttaaaaa agatatcat ttgaaagttc tcagagttgt acatatgttt cacagtacag gatctgtaca taaaagtttc ttctctaaac cattcaccaa gagccaatat ctaggcattt tcttggtagc acaaatttc ttattgctta gaaaattgtc ctccttgtta tttctgtttg taagacttaa gtgagttagg tctttaagga aagcaacgct cctctgaaat gcttgtcttt tatgctggga ggtgaccata gggctctgct ttaaaggat ttgagtgtctc aaaggccaga gtcacaggaa ggacttctc cagggagatt agtggtgatg ttaaaatgac ctcatgtcac tcttgtcac ggttttgttg agttttcact cttctaatgc aagggtctca cactgtgaac cacttaggat gtgatcactt tcaggtggc aggagagag aaggatctca cactgtgaac cacttaggat gtgatcactt tcaggtggc aggagagag aaggatctca cactgtgaac cacttaggat gtgatcactt tcaggtggc aggagatgtg aagggtctca cactgtgaac cacttaggat gtgatcactt tcaggtggc aggaatgttg aagggtctca cactgtgaac cacttaggat gtgatcactt tcaggtggc aggaatgttg aaggttcta tcaggttga tttaaaaaaa gtatctattt	149 197 257 317 377 437 497 557 617 677 737 797 857
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt cacagtacag gatctgtaca taaaagttc ttcatagat cattagttt cacagtacag gatctgtaca taaaagttc ttattgctta gaaaattgtc ctccttgta ttctgtttg taagacttaa gtgagtagg tcttaaagga aagcaacgct ctcttgaat gcttgtctt tatgctgga ggtgaccata gggctctgct tttaaaggat gagaggagag	149 197 257 317 377 437 497 557 617 677 737 797 857 917
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag aga ggt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tggaatgttt tggctcagtt cattaaaaa agatatctat ttgaaagttc tcaggagtatt tcttctgttg taagacttaa gtaggtatagt ttttcttgtta cacagtacag gatctgtaca acaaattttc ttattgctta gaaaattgtc tttctgtttg taagacttaa gtgagtagg tctttaagga aagcaacgct cccttgtta tttctgtttg taagacttaa gtgagtagg tctttaagga aagcaacgct cccttgtaa aaggccaga gtcacaggaa ggactcttc cagggagatt tcttaaagaa ttggctgcttc aagggccaga gtcacaggaa ggactcttc cagggagatt tctaaagaa ttggctgttc aagggccaga cccattagga ggtgaccata gggctctgct tctaaaagaa tcacattgcca aagggtctca cacttaggat gtatcactt tcaggtggc agaaggagag ttaaaatgac cccatgtgaac cacttaggat gtatcactt tcaggtggcc agaagtttg aatgttttca cagtacagga gtcacaagaa gaattttct ttataaaaaa atatctatt gaaagttctc aaggattgtca aatgtctttg gccagttca tcttgaccac aaagttctt tcaggtggcc agaattttc attgttcac cagtacagga cacttaagaa aatttctt tcagtagaca cacttcaagaa accactacta aaggatttac tctgacaca aaagttctt tcagaagccc tctccaaagaa accactacta aaggatttac tctgacaca aaagttctt tcagaagtcac tctcaaaaca accactactagaa agaatttct tctgacaca aaagttctt tcagaagtacaca tcctaacaca accactactagaa agaatttaa tctgacacaa aaagttcct tcaaaacca tccaacaaga accactacta aaggatttac tctgacacaa aaagttcct tccaaaacca tccaacaaga accactacta aagacttaa aaagttcct tccaacacaa aaagttcct tccaacacacaa aaagttcct tccaacacacaacaca	149 197 257 317 377 437 497 557 617 677 737 797 857 917
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag aga aga ggt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgttt cacagtacag gatctgtaca taaaagttc tttcctaaac gagccaatat ctaggcattt tcttggtag cacaatttc ttattgctta gaaaattgtc tttctgttt taagacttaa gtgagtagg tctttaaggaaggaggaggaggaggaggaggaggaggaggag	149 197 257 317 377 437 497 557 617 677 737 797 857 917 977
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg agg gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcaggatgtt tcacagtacag gatctgtaca taaaagtttc tttectaaac cattcaccaa gagccaatat ctaggcattt tcttggtagc acaaatttc ttattgcta gagaaattgtc tctcttgtta tttctgtttg taagacttaa gggactatc tctttaaga aagcaacgct cctctgaat gctgcttt tatgctgga gggaccata gggctctgct tataaagaa ggccaga gtcacaggaa ggacttctc caggagatt tctttaagga aggccatca aagggccaga gtcacaggaa ggacttctc caggagaatt tcaggagatg ttaaaaatgac ctcatgtcct tcttgccac ggttttgttg gagagagagg ctcaattct ggccagttca tcttaggat gagatcactt caggagatt tcaggagatg aatgtctttg gccagttca tcttgccac aatgttctt tcaggagat tccaaggaag gccaatacc aggcatttc tcttgccac aatgttctt tcaggagat tctcaacaca acacttagga gccaatacc aggcatttc tcttgccac aatgttctt tcaggagaga gccaatact tcctgtaac atatgtttca cagtacaga tctgtacat aaagtttctt tcctaaacca ttcaccaaga gccaatacc aggcatttc tctgtacaa aaagtttctt tcctaaacca ttcaccaaga gccaatacc tctgtttgta agacttaagt tcgtacataag tctgtacact tcaaggagaa gcaacgctcc ccttgttat tctgtttgta agacttaagt tggccataagg tctggagagagagagagagagagagagagagagagagaga	149 197 257 317 377 437 497 557 617 677 737 797 857 917
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tggatgttt cacagtacag gatctgtaca taaaagttc ttttcctaaac cattcacaa gagccaatat ctctggttg taagacttaa gtgagtcag ttttctgtttt tatgcttggag ggtgaccata gttctttatgcttt tatgctgga ggtgaccata gggtctgct tttatgcttt tatgctgtga acaaattttc tttattgctta gaacaacgct cccttgaaat tggtgttctc aagggtctca cacttaggaa ggacttctc cagggagatt tcattaagaa ggacttctc cagggagatt taaaatgac ctcatgcac tcttgtcaca gggtttgtg aggtttgtg agggaggagag attaaaatgac cacttaggaa ggacttctc cagggagatt tcaaggatgag ggtgtttgtt cacgaggagat aagggaggagag attaaaatgac cacttaggaa cacttaggaa ggattttctc cagggagatt tcaagggtcaatat caggattttc tcttgaaaa aagttcttc tcaagggtcaatat cacttaggaa ggattttcac tcttctaatgga ggaggttttag ggtgatcactt tcaaggatggaggaggattaggtctaatatgttca cacttaggaa gagttttcact tcttaaaggaa gagttttcact tcttaaagga aagttttcac tcttcaacaa gagattttc tcttgaaacca aattttct taaggatggccagagttaggccagagttaggccagaatttc tcttgaaaacca aaagtttct tcctaaacca aaggttttc tcttgaaacca aaagtttct tcctaaacca aaagttctc aagactaacct tctgaaacca cacttaaga gagttaaggc tcctaaacca aaagtttct tcctaaacca aaagttctt tcctaaacca aaagtttct tcctaaacca aaagttctc aagagtagaaccacaccac	149 197 257 317 377 437 497 557 617 677 737 797 857 917 977 1037 1097 1157 1217
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro 15 20 25 ggg aga tta gtg gtg atg gag agg agg gtt aaa atg acc tca tgt cct Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro 30 35 40 tct tgt cca cgg ttt tgt tgagttttca ctcttctaat gcaagggtct Ser Cys Pro Arg Phe Cys 45 cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt tggctcagtt catttaaaaa agatatctat ttgaaagttc tcaggatgtt tcacagtacag gatctgtaca taaaagtttc tttectaaac cattcaccaa gagccaatat ctaggcattt tcttggtagc acaaatttc ttattgcta gagaaattgtc tctcttgtta tttctgtttg taagacttaa gggactatc tctttaaga aagcaacgct cctctgaat gctgcttt tatgctgga gggaccata gggctctgct tataaagaa ggccaga gtcacaggaa ggacttctc caggagatt tctttaagga aggccatca aagggccaga gtcacaggaa ggacttctc caggagaatt tcaggagatg ttaaaaatgac ctcatgtcct tcttgccac ggttttgttg gagagagagg ctcaattct ggccagttca tcttaggat gagatcactt caggagatt tcaggagatg aatgtctttg gccagttca tcttgccac aatgttctt tcaggagat tccaaggaag gccaatacc aggcatttc tcttgccac aatgttctt tcaggagat tctcaacaca acacttagga gccaatacc aggcatttc tcttgccac aatgttctt tcaggagaga gccaatact tcctgtaac atatgtttca cagtacaga tctgtacat aaagtttctt tcctaaacca ttcaccaaga gccaatacc aggcatttc tctgtacaa aaagtttctt tcctaaacca ttcaccaaga gccaatacc tctgtttgta agacttaagt tcgtacataag tctgtacact tcaaggagaa gcaacgctcc ccttgttat tctgtttgta agacttaagt tggccataagg tctggagagagagagagagagagagagagagagagagaga	149 197 257 317 377 437 497 557 617 677 737 797 857 917 977 1037 1097 1157

caccaagagc	caatatctag	gcattttctt tgtttgtaag	ggtagcacaa acttaagtga	gttaggtctt	tgcttagaaa taaggaaagc tctgctttta	1397 1457 1517

<210> 76 <211> 526 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 22..318 <221> sig_peptide <222> 22..93 <223> Von Heijne matrix score 4.6 seq FFIFCSLNTLLLG/GV <221> polyA_signal <222> 497..502 <221> polyA_site <222> 516..526 <400> 76 ctgcctgctg cttgctgcac c atg aag tct gcc aag ctg gga ttt ctt cta 51 Met Lys Ser Ala Lys Leu Gly Phe Leu Leu -20 99 aga tto tto ato tto tgo toa ttg aat acc ctg tta ttg ggt ggt gtt Arg Phe Phe Ile Phe Cys Ser Leu Asn Thr Leu Leu Gly Gly Val -5 -10 aat aaa att gcg gag aag ata tgt gga gac ctc aaa gat ccc tgc aaa Asn Lys Ile Ala Glu Lys Ile Cys Gly Asp Leu Lys Asp Pro Cys Lys 15 10 195 ttg gac atg aat ttt gga agc tgc tat gaa gtt cac ttt aga tat ttc Leu Asp Met Asn Phe Gly Ser Cys Tyr Glu Val His Phe Arg Tyr Phe 25 tac aac aga acc tcc aaa aga tgt gaa act ttt gtc ttc tcc ggc tgt 243 Tyr Asn Arg Thr Ser Lys Arg Cys Glu Thr Phe Val Phe Ser Gly Cys 45 40 aat ggc aac ctt aac aac ttc aag ctt aaa ata gaa cgt gaa gta gcc 291 Asn Gly Asn Leu Asn Asn Phe Lys Leu Lys Ile Glu Arg Glu Val Ala 60 55 338 tgt gtt gca aaa tac aaa cca ccg agg tgagaggatg tgaactcatg Cys Val Ala Lys Tyr Lys Pro Pro Arg 70 aagttgtctg ctgcaccatc cgaaataaag acacaagaaa attcagactg attttgaaat 398 ctttgtaata tttccataat gctttaagct tccatatgtt tgctattttc ctgaccctag 458 518 ttttgtcttt cctggaaatt aactgtatga tcattagaat gaaagagtct ttctgtcaaa 526 aaaaaaa

<210> 77 <211> 352 <212> DNA <213> Homo sapiens

<220>	>															
<221:	> CD	S														
<222:	> 8.	.292												,		
<221:	> si	g_pe	ptid	e												
<222	> 8.	.118														
<223	> Vo	n He	ijne	mat	rix											
	sc	ore	5.6													
	se	a WI	LLDA	LLRI	GDT/	KK										
		•														
<221:	> pc	lyA	sign	al												
<222:	_		. –													
<221	> pc	lyA	site	:												
<222	_		-													
<400	> 77	,														
ctga	gat	atq	qca	agt	ccc	gct	gta	aac	agg	tgg	aaa	agg	cca	agg	ttg	49
J ,	J	Met	Ãla	Ser	Pro	Āla	Val	Asn	Arg	Trp	Lys	Arg	Pro	Arg	Leu	
				-35					-30					-25		
aag	ccq	ata	tgg	cca	cgg	cgc	ttg	gaa	tcc	tgg	ttg	ttg	ctg	gat	gct	97
Lys	Pro	Val	Trp	Pro	Arg	Arg	Leu	Glu	Ser	Trp	Leu	Leu	Leu	Asp	Ala	
			-20		•	_		-15					-10			
ctt	ttq	cqa	tta	gga	gat	acc	aaa	aaa	aag	cga	cag	cct	gaa	gca	gcc	145
Leu	Leu	Arq	Leu	Gly	Asp	Thr	Lys	Lys	Lys	Arg	Gln	Pro	Glu	Ala	Ala	
		-5		•	-		1				5					
aca	aaa	tcc	tat	qtt	aga	agc	agc	tgt	999	ggt	CCC	agt	gga	gat	999	193
Thr	Lvs	Ser	Cys	Val	Arg	Ser	Ser	Cys	Gly	Gly	Pro	Ser	Gly	Asp	GIÀ	
10					15					20					25	
cct	ccc	cca	tqc	ctc	cag	cag	cct	gac	cct	cgt	gcc	ctg	tct	cag	gcg	241
Pro	Pro	Pro	Cys	Leu	Gln	Gln	Pro	Asp	Pro	Arg	Ala	Leu	Ser	Gln	Ala	
				30					35					40		
ttc	tct	aga	tcc	ttt	cct	ctg	ttt	ccc	tct	ctc	gct	ggc	aaa	agt	atg	289
Phe	Ser	Ara	Ser	Phe	Pro	Leu	Phe	Pro	Ser	Leu	Ala	Gly	Lys	Ser	Met	
		5	45					50					55			
atc	taat	ttqaa		aaga	ctqaa	ag g	atca	ataa	a ca	gcca	tctg	ccc	cttc	aaa		342
Ile		5 1			5	ر ر				_						
aaaa	aaa	aaa														352

<221> polyA_site < <222> 522..542

.400. 79	
<400> 78 cacgacctgt gggcc atg atg cta ccc caa tgg ctg ctg ctg ctg ttc ctt Met Met Leu Pro Gln Trp Leu Leu Leu Phe Leu	51
-20 -15	
ctc ttc ttc ttt ctc ttc ctc ctc acc agg ggc tca ctt tct cca aca	99
Leu Phe Phe Leu Phe Leu Leu Thr Arg Gly Ser Leu Ser Pro Thr -10 -5 1 5	
aaa tat aac ctt ttg gag ctc aag gag tct tgc atc cgg aac cag gac	147
Lys Tyr Asn Leu Leu Glu Leu Lys Glu Ser Cys Ile Arg Asn Gln Asp 10 15 20	
tgc gag act ggc tgc tgc caa cgt gct cca gac aat tgc gag tcg cac	195
Cys Glu Thr Gly Cys Cys Gln Arg Ala Pro Asp Asn Cys Glu Ser His 25 30 35	
tgc gcg gag aag ggg tcc gag ggc agt ctg tgt caa acg cag gtg ttc	243
Cys Ala Glu Lys Gly Ser Glu Gly Ser Leu Cys Gln Thr Gln Val Phe 40 45 50	
ttt ggc caa tat aga gcg tgt ccc tgc ctg cgg aac ctg act tgt ata	291
Phe Gly Gln Tyr Arg Ala Cys Pro Cys Leu Arg Asn Leu Thr Cys Ile 55 60 65	
tat toa aag aat gag aaa tgg ott ago ato goo tat ggo ogt tgt cag	339
Tyr Ser Lys Asn Glu Lys Trp Leu Ser Ile Ala Tyr Gly Arg Cys Gln 70 75 80 85	200
aaa att gga agg cag aag ttg gct aag aaa atg ttc ttc tagtgctccc	388
Lys Ile Gly Arg Gln Lys Leu Ala Lys Lys Met Phe Phe 90 95	
toottottgo tgoctootoo tootooacot gototootoo otaccoagag ctotgtgtto	448
accetgitee ceagageete caccatgagt ggagggaagt ggggagtgat tgaaataaag	508
agctttttca atgaaaaaaa aaaaaaaaaa aaaa	542
<210> 79	
<211> 233	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 57233	
<400> 79	
gcaaaaccaa aaccagcacc gatcccgaca tagatcagtg acgtcttttt cttcag atg Met	59
ato ota tgt tto ott ott oot oat oat ogt ott oag gaa goo aga cag	107
Ile Leu Cys Phe Leu Leu Pro His His Arg Leu Gln Glu Ala Arg Gln	
5 10 15	
att caa gta ttg aag atg ctg cca agg gaa aaa tta aga aga gaa	155
Ile Gln Val Leu Lys Met Leu Pro Arg Glu Lys Leu Arg Arg Arg Glu 20 25 30	
gag aga aaa caa ata aat ggg aaa aaa gaa agg aca aaa tat gaa aca	203
Glu Arg Lys Gln Ile Asn Gly Lys Lys Glu Arg Thr Lys Tyr Glu Thr 35 40 45	
cca aga aaa aga gaa gga aaa aaa aaa aaa	233
Pro Arg Lys Arg Glu Gly Lys Lys Lys	
50 55	

<210> 80 <211> 660 <212> DNA

```
<213> Homo sapiens
<220>
<221> CDS
<222> 83..340
<221> sig_peptide
<222> 83..124
<223> Von Heijne matrix
     score 7.5
      seq VALNLILVPCCAA/WC
<221> polyA_signal
<222> 573..578
<221> polyA_site
<222> 607..660
<400> 80
gaatttgtaa aacttctgct cgtttacact gcacattgaa tacaggtaac taattggaag
                                                                      60
gagaggggag atcactcttt tg atg gtg gcc ctg aac ctc att ctg gtt ccc
                                                                      112
                         Met Val Ala Leu Asn Leu Ile Leu Val Pro
tgc tgc gct gct tgg tgt gac cca cgg agg atc cac tcc cag gat gac
                                                                      160
Cys Cys Ala Ala Trp Cys Asp Pro Arg Arg Ile His Ser Gln Asp Asp
gtg ccc cgt agc tct gct gct gat act ggg tct gcg atg cag cgg cgt
Val Pro Arg Ser Ser Ala Ala Asp Thr Gly Ser Ala Met Gln Arg Arg
                                                25
                            20
gag gcc tgg gct ggt tgg aga agg tca caa ccc ttc tct gtt ggt ctg
Glu Ala Trp Ala Gly Trp Arg Arg Ser Gln Pro Phe Ser Val Gly Leu
                        35
                                                                      304
cct tct gct gaa aga ctc gag aac caa cca ggg aag ctg tcc tgg agg
Pro Ser Ala Glu Arg Leu Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg
                                     55
                                                                      350
tee etg gte gga gag gga tat aga ate tgt gae ete tgacaactgt
Ser Leu Val Gly Glu Gly Tyr Arg Ile Cys Asp Leu
                                    70
                65
gaagccaccc tgggctacag aaaccacagt cttcccagca attattacaa ttcttgaatt
                                                                      410
ccttggggat tttttactgc cctttcaaag cacttaagtg ttagatctaa cgtgttccag
                                                                      470
tgtctgtctg aggtgactta aaaaatcaga acaaaacttc tattatccag agtcatggga
                                                                      530
                                                                      590
gagtacaccc tttccaggaa taatgttttg ggaaacactg aaatgaaatc ttcccagtat
tataaattgt gtatttaaaa aaagaaactt ttctgaatgc ctacctggcg gtgtatacca
                                                                      650
                                                                      660
ggcagtgtgc
```

<210> 81
<211> 605
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 47..541

<221> sig_peptide
<222> 47..220
<223> Von Heijne matrix
score 5.4
seq QLLDSVLWLGALG/LT

<221> polyA_site <222> 597..605

<pre><400> 81 aaagtgggag gagcactagg tettecegte acetecacet etetee atg ace egg</pre>	55
Met Thr Arg	
ctc tgc tta ccc aga ccc gaa gca cgt gag gat ccg atc cca gtt cct	103
Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro Ile Pro Val Pro	
-55 -50 -45 -40	
cca agg ggc ctg ggt gct ggg gag ggg tca ggt agt cca gtg cgt cca	151
Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser Pro Val Arg Pro	
-35 -30 -25	
cot gta too acc tgg ggc cot ago tgg gcc cag oto otg gac agt gto	199
Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu Leu Asp Ser Val	
-20 -15 -10	
cta tgg ctg ggg gca cta gga ctg aca atc cag gca gtc ttt tcc acc	247
Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala Val Phe Ser Thr	
-5 1 5	
act ggc cca gcc ctg ctg ctt ctg gtc agc ttc ctc acc ttt gac	295
Thr Gly Pro Ala Leu Leu Leu Leu Val Ser Phe Leu Thr Phe Asp	
10 15 20 25	
ctg ctc cat agg ccc gca ggt cac act ctg cca cag cgc aaa ctt ctc	343
Leu Leu His Arg Pro Ala Gly His Thr Leu Pro Gln Arg Lys Leu Leu	
30 35 40	391
acc agg ggc cag agt cag ggg gcc ggt gaa ggt cct gga cag cag gag	231
Thr Arg Gly Gln Ser Gln Gly Ala Gly Glu Gly Pro Gly Gln Gln Glu	
45	439
get cta etc etg caa atg ggt aca gte tea gga caa ett age etc eag	
Ala Leu Leu Gln Met Gly Thr Val Ser Gly Gln Leu Ser Leu Gln 60 65 70	
gac gca ctg ctg ctg ctc atg ggg ctg ggc ccg ctc ctg aga gcc	487
Asp Ala Leu Leu Leu Leu Met Gly Leu Gly Pro Leu Leu Arg Ala	
75 80 85	
tgt ggc atg ccc ttg acc ctg ctt ggc ctg gct ttc tgc ctc cat cct	535
Cys Gly Met Pro Leu Thr Leu Leu Gly Leu Ala Phe Cys Leu His Pro	
90 95 100 105	
tgg gcc tgagagcccc tccccacaac tcagtgtcct tcaaatatac aatgaccacc	591
Trp Ala	
cttcttcaaa aaaa	605

<210> 82

<211> 396

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 46..285

<221> sig_peptide

<222> 46..150

<223> Von Heijne matrix score 3.6 seq LEPGLSSSAACNG/KE

<221> polyA_signal

<222> 364..369

<221> polyA_site <222> 385..396

<pre><400> 82 cctctacagg aatcagactc agcctctttt ggttttcagt gaagt atg cct ttt caa</pre>	. 57
ttt gga acc cag cca agg agg ttt cca gtg gaa gga gga gat tct tca Phe Gly Thr Gln Pro Arg Arg Phe Pro Val Glu Gly Gly Asp Ser Ser -30 -25 -20	105
att gag ctg gaa cct ggg ctg agc tcc agt gct gcc tgt aat ggg aag Ile Glu Leu Glu Pro Gly Leu Ser Ser Ala Ala Cys Asn Gly Lys -15 -10 -5 1	153
gag atg tca cca acc agg caa ctc cgg agg tgc cct gga agt cat tgc Glu Met Ser Pro Thr Arg Gln Leu Arg Arg Cys Pro Gly Ser His Cys 5 10 15	201
ctg aca ata act gat gtt ccc gtc act gtt tat gca aca acg aga aag Leu Thr Ile Thr Asp Val Pro Val Thr Val Tyr Ala Thr Thr Arg Lys 20 25 30	249
cca cct gca caa agc agc aag gaa atg cat cct aaa tagcaccatt Pro Pro Ala Gln Ser Ser Lys Glu Met His Pro Lys 35 40 45	295
aagtetttig teaaggtetg actaggteaa gggtaatgga eeagtateat etggtgatet ggtaaacaaa taaaagtggt ggeaeettea aaaaaaaaa a	355 396
<210> 83	
<pre><210> 63 <211> 432 <212> DNA <213> Homo sapiens</pre>	
<220>	
<2205 <221> CDS <222> 22240	
<221> sig_peptide <222> 2284 <223> Von Heijne matrix score 12 seq VLVLCVLLLQAQG/GY	
<221> polyA_signal <222> 397402	
<221> polyA_site <222> 421432	
<pre><400> 83 gctcacgctc tggtcagagt t atg gca ccc cag act ctg ctg cct gtc ctg</pre>	51
gtt ctc tgt gtg ctg ctg ctg cag gcc cag gga gga tac cgt gac aag Val Leu Cys Val Leu Leu Leu Gln Ala Gln Gly Gly Tyr Arg Asp Lys -10 -5 1 5	99
atg agg atg cag aga atc aag gtc tgt gag aag cga ccc agc ata gat Met Arg Met Gln Arg Ile Lys Val Cys Glu Lys Arg Pro Ser Ile Asp 10 15 20	147
cta tgc atc cac tgt tca tgt ttc caa aag tgt gaa aca aat aag Leu Cys Ile His His Cys Ser Cys Phe Gln Lys Cys Glu Thr Asn Lys 25 30 35	195
ata tgc tgt tca gcc ttc tgt ggg aac att tgt atg agc atc cta Ile Cys Cys Ser Ala Phe Cys Gly Asn Ile Cys Met Ser Ile Leu 40 45 50	240

tgagtgggag agtgggctgg gatgtgcatc ctgctccctg aaccettcca tccgagactg tgcccacatc cgaagcacaa ggacatcaaa tcatcagcac aagaacatca acaggaatgc caccetecce agtgtctgaa etceetgtee etgtcaaatg aaccagaaca aatgcccatg aaaaaaaaaa aa 432	0
<210> 84 <211> 420 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 89382	
<221> polyA_site <222> 408420	
<pre><400> 84 gcttgcctga ccccatgtc gcctctgtag gtagaagaag tatgtcttcc tggaccccct ggctggtgct gtaacaaaga cccatgtg atg ctg ggg gca gag aca gag gag Met Leu Gly Ala Glu Thr Glu Glu 1</pre> 1	
aag ctg ttt gat gcc ccc ttg tcc atc agc aag aga gag cag ctg gaa 160 Lys Leu Phe Asp Ala Pro Leu Ser Ile Ser Lys Arg Glu Gln Leu Glu 10 15 20)
cag cag gtc cca gag aac tac ttc tat gtg cca gac ctg ggc cag gtg Gln Gln Val Pro Glu Asn Tyr Phe Tyr Val Pro Asp Leu Gly Gln Val 25 30 35 40	3
cct gag att gat gtt cca tcc tac ctg cct gac ctg ccc ggc att gcc 256 Pro Glu Ile Asp Val Pro Ser Tyr Leu Pro Asp Leu Pro Gly Ile Ala 45 50 55	5
aac gac ctc atg tac att gcc gac ctg ggc ccc ggc att gcc ccc tct 304 Asn Asp Leu Met Tyr Ile Ala Asp Leu Gly Pro Gly Ile Ala Pro Ser 60 65 70	Ł
gcc cct ggc acc att cca gaa ctg ccc acc ttc cac act gag gta gcc 352 Ala Pro Gly Thr Ile Pro Glu Leu Pro Thr Phe His Thr Glu Val Ala 75 80 85	2
gag cct ctc aag acc tac aag atg ggg tac taacagcacc accaccgccc 402 Glu Pro Leu Lys Thr Tyr Lys Met Gly Tyr 90 95	2
CCACCAAAA AAAAAAA 420)
<210> 85 <211> 501 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 80415	
<221> sig_peptide <222> 80142 <223> Von Heijne matrix	

<221> polyA_signal

<223> Von Heijne matrix score 5.4

seq TFCLIFGLGAVWG/LG

<222> 471476	
<221> polyA_site <222> 488501	
<400> 85 cccgcttgat tccaagaacc tcttcgatat ttattttat ttttaaagag ggagacgatg gactgagctg atccgcacc atg gag tct cgg gtc tta ctg aga aca ttc tgt Met Glu Ser Arg Val Leu Leu Arg Thr Phe Cys -20 -15	60 112
ttg atc ttc ggt ctc-gga gca gtt tgg ggg ctt ggt gtg gac cct tcc Leu Ile Phe Gly Leu Gly Ala Val Trp Gly Leu Gly Val Asp Pro Ser	160
cta cag att gac gtc tta aca gag tta gaa ctt ggg gag tcc acg acc Leu Gln Ile Asp Val Leu Thr Glu Leu Glu Leu Gly Glu Ser Thr Thr	208
gga gtg cgt cag gtc ccg ggg ctg cat aat ggg acg aaa gcc ttt ctc Gly Val Arg Gln Val Pro Gly Leu His Asn Gly Thr Lys Ala Phe Leu 25 30 35	256
ttt caa gat act ccc aga agc ata aaa gca tcc act gct aca gct gaa Phe Gln Asp Thr Pro Arg Ser Ile Lys Ala Ser Thr Ala Thr Ala Glu 40 45 50	304
cag ttt ttt cag aag ctg aga aat aaa cat gaa ttt act att ttg gtg Gln Phe Phe Gln Lys Leu Arg Asn Lys His Glu Phe Thr Ile Leu Val 55 60 65 70	352
acc cta aaa cag acc cac tta aat tca gga gtt att ctc tca att cac Thr Leu Lys Gln Thr His Leu Asn Ser Gly Val Ile Leu Ser Ile His 75 80 85	400
cac ttg gat cac agg taaatgtggt tgctggagtt tcctgtgttt tcattatatg His Leu Asp His Arg	455
90 tggttaaatg aatatattaa agagaagtaa acaaaaaaaa aaaaaa	501
<210> 86 <211> 454 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 152361	
<221> sig_peptide <222> 152283 <223> Von Heijne matrix score 4.7 seq FLLSLSLITYCFW/DP	
<pre><400> 86 gacattttac ttttttctgt taacgcttac cctagaaatt agaaatgaca ccacgtattc ttagcgaagt ccagttttca gcattttgtc cttattggac aatagcaagg atattagaac gtgttggttc cgcgtgcttc cgtcttgagt t atg tgc tgc tat tgt cgg ata</pre>	60 120 172
ttt tgt ctt aga tgt acg tac ttt cct gtt cat tgt ggt atg tgt aat Phe Cys Leu Arg Cys Thr Tyr Phe Pro Val His Cys Gly Met Cys Asn -35 -30 -25	220
ttg cgt tac ttt gaa ttt tcc acg ttt tta ctt tct ttg tct ctc atc Leu Arg Tyr Phe Glu Phe Ser Thr Phe Leu Leu Ser Leu Ser Leu Ile -20 -15 -10	268

act tac tgc ttt tgg gac ccc ccc cat cgg ggt tca cat tcc ctc tcc	316
Thr Tyr Cys Phe Trp Asp Pro Pro His Arg Gly Ser His Ser Leu Ser	
_5 1 5 10 · · · · · · · · · · · · · · · · · ·	
cta gag cac act ccc ttg gat ttc ctc gag tgg ggt ctg ctg cgg	361
Leu Glu His Thr Pro Leu Asp Phe Leu Glu Trp Gly Leu Leu Arg	
15 20 25	421
tgaagctttc ccattttatg tgcagattat tttcagaggg tatatagaat tcaggcagct	454
gtttcgttgt agcacattaa aaatattttc ccc	454
<210> 87	
<211> 1272	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 32307	
<pre><221> sig_peptide <222> 3270</pre>	
<pre><222> 32/0 <223> Von Heijne matrix</pre>	
score 4.2	
seq MLFSLSLLSNLNQ/IG	
<221> polyA signal	
<222> 12401245	
<221> polyA_site	
<222> 12611272	
.400. 07	
<400> 87 Strangetter agetteergag c atg ctg fft tet etc age ett	52
gtcaggttgc accgecettt ggttecegag c atg etg ttt tet ete age ett	52
<pre><400> 87 gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt</pre>	52
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac	52
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His	
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 10	100
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa	
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln	100
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25	100
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac	100
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His	100
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40	100
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa	100 148 196
gtcaggttgc accgccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40	100 148 196
gtcaggttgc accgccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 50 55 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca	100 148 196
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys	100 148 196 244
gtcaggttgc accgcccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65	100 148 196 244 292
gtcaggttgc acceccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 Ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5	100 148 196 244
gtcaggttgc accepecttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5	100 148 196 244 292
gtcaggttgc accepted t ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65 70 cca ttt cta gct tgc tgaaacccaa aactaatctc caagaaggag aagcttctct Pro Phe Leu Ala Cys 75	100 148 196 244 292
gtcaggttgc accecettt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 50 55 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65 70 cca ttt cta gct tgc tgaaacccaa aactaatctc caaggaaggag aagcttctct Pro Phe Leu Ala Cys 75 cgcaggccgga gcaggtccct ttctagagat aggagaagag agagatcgct gtctcggag	100 148 196 244 292 347
gtcaggttgc accgccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 50 55 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65 70 cca ttt cta gct tgc tgaaacccaa aactaatctc caagaaggag agagttcctt Pro Phe Leu Ala Cys 75 cgcagccgga gcaggtccct ttctagagat aggagaagag agagatcgct gtctcggag agaaatcaca agccgtcccg atccttctct aggtctcgta gtcgatttag gtcaaatgaa	100 148 196 244 292 347 407 467
gtcaggttgc acceccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 50 55 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65 70 cca ttt cta gct tgc tgaaacccaa aactaatctc caagaaggag aggatcgct gtctcgggag agaaatcaca agccgtcccg atccttctct aggtctcgta gtcgatttag gtcaaatgaa aggaaatcaca agacgtcccg atccttctct aggtctcgta gtcgatttag gtcaaatgaa aggaaatcaca agacgttccc atctctctct aggtctcaagg aaattacttc atttgacagg	100 148 196 244 292 347 407 467 527
gtcaggttgc acceccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 Ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 Caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 Cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65 70 Cca ttt cta gct tgc tgaaacccaa aactaatctc caagaaggag aagcttctct Pro Phe Leu Ala Cys 75 Cgcagccgga gcaggtccct ttctagagat aggagaagag agagatcgct gtctcgggag agaaatcaca agccgtcccg atccttctct aggtctcgta gtcgatttag gtcaaatgaa aggaaataga agacagtttg caagagaagt ggtgtacaagg aaattacttc atttgacagg agtaattacta gaaaattcaa gttttgtttg agacttcata agcttggtgc atttttaaga	100 148 196 244 292 347 407 467
gtcaggttgc accegccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aca gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 50 55 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65 70 cca ttt cta gct tgc tgaaacccaa aactaatctc caagaaggag aagattcct Pro Phe Leu Ala Cys 75 cgcagccgga gcaggtccct ttctagagat aggagaagag agagatcgct gtccgggag agaaatcaca agccgtcccg atccttctct aggtctcgta gtcgatttag gtcaaatgaa aggaaatagaa gacaagtttg caagaagat ggtgtacagg aactaatacttc atttgacagg agtatgtaca gaaaatcaa gttttgtttg agacttcata agcttggtgc atttttaaga tgttttaagct gttcaaacc gtttgttct tgaaaccaaaagt gacacaaaagt gtaaattcct	100 148 196 244 292 347 407 467 527 587
### Second Company of the company of	100 148 196 244 292 347 467 527 587 647
gtcaggttgc accegccttt ggttcccgag c atg ctg ttt tct ctc agc ctt Met Leu Phe Ser Leu Ser Leu -10 ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His -5 1 5 10 att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln 15 20 25 caa cta cag cag cag cct tcg gct aac aaa aca gca gga aaa atc cac Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His 30 35 40 aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys 45 50 55 cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser 60 65 70 cca ttt cta gct tgc tgaaacccaa aactaatctc caagaaggag aagattcct Pro Phe Leu Ala Cys 75 cgcagccgga gcaggtccct ttctagagat aggagaagag agagatcgct gtccgggag agaaatcaca agccgtcccg atccttctct aggtctcgta gtcgatttag gtcaaatgaa aggaaatagaa gacaagtttg caagaagat ggtgtacagg aactaatacttc atttgacagg agtatgtaca gaaaatcaa gttttgtttg agacttcata agcttggtgc atttttaaga tgttttaagct gttcaaacc gtttgttct tgaaaccaaaagt gacacaaaagt gtaaattcct	100 148 196 244 292 347 467 527 587 647 707

-- -

		_	· • ·	_ _ ·				+	-+~~	79 F	ateor	rct a	ca c	ctat	agttg	887
22022	acac	rt ct	ttaa	aaaa	c ta	ctate	gaaa	caca	aggc	cat (cagg	gaaa	ac g	aaat	gctgc	947
actat	taaa	it ta	agage	sttti	t ta	aaaaa	atcc	aac	tctca	atc (ctgg	gcaga	ag g	ttgc	ctagt	1007 1067
ttgat	ttaco	a ta	atato	ggata	a cat	tggci	tgtt	cgt	gacai	ttc	tttai	tgtg	ca a	attt	gtgat	1127
ttcaa	aaaat	a to	ccta	ccagi	t tt	aagg	gtac	att	gtaga	agc ·	cgaa	cttt	ga g	ttac	tgtgc	1187 1247
aaga tttg	gttac	t ti	ccat; attaa	gctg	a aa	aaa	Lddl	acg		gry .	agaa		כ כ	9400	aaagt	1272
	,															
<210																
	> 804 > DNA															
	> Hor		apie:	ns												
<220	>															
	> CDS		. .													
<222	> 114	± /	34													
	> sig > 114			е												
	> Aoi			mat	rix											
		ore	-	CHEF	cos/	ממ										
		_			CQ5/											
	> po:		_	al												
-221	> po	Δν:1	gite								•					
	> 79	-														
<400	> 88															
ccaa	> 88	ag g	aaga	gtct	g aa	gago	agco	agt	gttt	.cgg	cttg	tgcc	ct g	tata	icttga itg	60 116
ccaa agct	cacc gcca	aa c	aagt	acgg	jt ag	ittet	gaaa	a ato	caga	atg	gctt	gatg	וככ נ	ac a N	iet Iet	116
ccaa agct	cacc gcca att	aa c tta	aagt caa	acgg	gt ag	act	gaaa	a ato gtg	caga gat	gat	gctt	gatg att	caa	ac a N gca	let att	
ccaa agct cac His	cacc gcca att Ile -40	aa c tta Leu	caa Gln	acgg ctg Leu	rt ag ctt Leu	act Thr	gaaa aca Thr	gtg Val	caga gat Asp	gat Asp	gga Gly -30	att Ile	caa Gln	gca Ala	Met att Ile	116 164
ccaa agct cac His	cacc gcca att Ile -40	aa c tta Leu	caa Gln	ctg Leu	ctt Leu	act Thr -35	aca Thr	gtg Val	gat Asp att	gat Asp	gga Gly -30	gatg att Ile tta	caa Gln	gca Ala	Met att Ile gac	116
ccaa agct cac His gta Val	cacc gcca att Ile -40 cat His	aa c tta Leu tgt Cys	caa Gln cct Pro	ctg Leu gac Asp	ctt Leu act Thr	act Thr -35 gga Gly	aca Thr aaa Lys	gtg Val gac Asp	gat Asp att Ile	gat Asp tgg Trp -15	gga Gly -30 aat Asn	att Ile tta Leu	caa Gln ctt Leu	gca Ala ttt Phe	Met att Ile gac Asp	116 164 212
ccaa agct cac His gta Val -25	cacc gcca att Ile -40 cat His	aa c tta Leu tgt Cys	caa Gln cct Pro	ctg Leu gac Asp	ctt Leu act Thr	act Thr -35 gga Gly	aca Thr aaa Lys	gtg Val gac Asp	gat Asp att Ile	gat Asp tgg Trp -15	gga Gly -30 aat Asn	att Ile tta Leu	caa Gln ctt Leu	gca Ala ttt Phe	Met att Ile gac Asp -10 ctt	116 164
ccaa agct cac His gta Val -25	cacc gcca att Ile -40 cat His	aa c tta Leu tgt Cys	caa Gln cct Pro	ctg Leu gac Asp gaa Glu	ctt Leu act Thr	act Thr -35 gga Gly	aca Thr aaa Lys	gtg Val gac Asp	gat Asp att Ile	gat Asp tgg Trp -15	gga Gly -30 aat Asn	att Ile tta Leu	caa Gln ctt Leu	gca Ala ttt Phe	Met att Ile gac Asp -10 ctt	116 164 212
ccaa agct cac His gta Val -25 ctg Leu	att Ile -40 cat His gtc Val	aa c tta Leu tgt Cys tgc Cys	caa Gln cct Pro cat His	ctg Leu gac Asp gaa Glu -5	ctt Leu act Thr -20 ttc Phe	act Thr -35 gga Gly tgc Cys	aca Thr aaa Lys cag Gln	gtg Val gac Asp tct Ser	gat Asp att Ile gat Asp l	gat Asp tgg Trp -15 gat Asp	gga Gly -30 aat Asn cca Pro	att Ile tta Leu ccc Pro	caa Gln ctt Leu atc Ile 5	gca Ala ttt Phe att Ile	Met att Ile gac Asp -10 ctt Leu	116 164 212
ccaa agct cac His gta Val -25 ctg Leu	att Ile -40 cat His gtc Val	tta Leu tgt Cys tgc Cys cag Gln	caa Gln cct Pro cat His	ctg Leu gac Asp gaa Glu -5	ctt Leu act Thr -20 ttc Phe	act Thr -35 gga Gly tgc Cys	aca Thr aaa Lys cag Gln gcc Ala	gtg Val gac Asp tct Ser	gat Asp att Ile gat Asp l	gat Asp tgg Trp -15 gat Asp ttt	gga Gly -30 aat Asn cca Pro	att Ile tta Leu ccc Pro	caa Gln ctt Leu atc Ile 5	gca Ala ttt Phe att Ile	Met att Ile gac Asp -10 ctt Leu	116 164 212 260
ccaa agct cac His gta Val -25 ctg Leu caa Gln	att Ile -40 cat His gtc Val gaa Glu	tta Leu tgt Cys tgc Cys cag Gln 10	caa Gln cct Pro cat His aaa Lys	ctg Leu gac Asp gaa Glu -5 aca Thr	ctt Leu act Thr -20 ttc Phe gtg Val	act Thr -35 gga Gly tgc Cys cta Leu	aca Thr aaa Lys cag Gln gcc Ala 15 caa	gac yal yac Asp tct Ser tct Ser gag	gat Asp att Ile gat Asp 1 gtt Val	gat Asp tgg Trp -15 gat Asp ttt Phe cta	gga Gly -30 aat Asn cca Pro tca Ser	att Ile tta Leu ccc Pro gtg Val 20 ata	caa Gln ctt Leu atc Ile 5 ttg Leu	gca Ala ttt Phe att Ile tct Ser	Met att Ile gac Asp -10 ctt Leu gcc Ala	116 164 212 260
ccaa agct cac His gta Val -25 ctg Leu caa Gln	caccegcca att Ile -40 cat His gtc Val gaa Glu tat Tyr	tta Leu tgt Cys tgc Cys cag Gln 10	caa Gln cct Pro cat His aaa Lys	ctg Leu gac Asp gaa Glu -5 aca Thr	ctt Leu act Thr -20 ttc Phe gtg Val	act Thr -35 gga Gly tgc Cys cta Leu gag Glu	aca Thr aaa Lys cag Gln gcc Ala 15 caa	gac yal yac Asp tct Ser tct Ser gag	gat Asp att Ile gat Asp 1 gtt Val	gat Asp tgg Trp -15 gat Asp ttt Phe cta	gga Gly -30 aat Asn cca Pro tca Ser	att Ile tta Leu ccc Pro gtg Val 20 ata	caa Gln ctt Leu atc Ile 5 ttg Leu	gca Ala ttt Phe att Ile tct Ser	Met att Ile gac Asp -10 ctt Leu gcc Ala	116 164 212 260 308
ccaa agct cac His gta Val -25 ctg Leu caa Gln atc	cacca gcca att Ile -40 cat His gtc Val gaa Glu tat Tyr	tta Leu tgt Cys tgc Cys cag Gln 10 gcc Ala	caa Gln cct Pro cat His aaa Lys tca Ser	ctg Leu gac Asp gaa Glu -5 aca Thr cag Gln	ctt Leu act Thr -20 ttc Phe gtg Val act Thr	act Thr -35 gga Gly tgc Cys cta Leu gag Glu 30 agc	aca Thr aaa Lys cag Gln gcc Ala 15 caa Gln	gtg Val gac Asp tct Ser tct Ser gag Glu	gat Asp att Ile gat Asp gtt Val tat Tyr	gatg gat Asp tgg-15 gat Phe cta Leu gtc	gga Gly -30 aat Asn cca Pro tca Ser aag Lys 35 tta	att Ile tta Leu ccc Pro gtg Val 20 ata Ile caa	caa Gln ctt Leu atc Ile 5 ttg Leu gaa Glu aat	gca Ala ttt Phe att Ile tct Ser aaa Lys	Met att Ile gac Asp -10 ctt Leu gcc Ala gta Val	116 164 212 260 308
ccaa agct cac His gta Val -25 ctg Leu caa Gln atc Ile	cacca gcca att Ile -40 cat His gtc Val gaa Glu tat T25	tta Leu tgt Cys tgc Cys cag Gln 10 gcc Ala	caa Gln cct Pro cat His aaa Lys tca Ser	ctg Leu gac Asp gaa Glu -5 aca Thr cag Gln	ctt Leu act Thr -20 ttc Phe gtg Val act Thr	act Thr -35 gga Gly tgc Cys cta Leu gag Glu 30 agc	aca Thr aaa Lys cag Gln gcc Ala 15 caa Gln	gtg Val gac Asp tct Ser tct Ser gag Glu	gat Asp att Ile gat Asp gtt Val tat Tyr	gat Asp tgg Trp -15 gat Asp ttt Phe cta Leu Val	gga Gly -30 aat Asn cca Pro tca Ser aag Lys 35 tta	att Ile tta Leu ccc Pro gtg Val 20 ata Ile caa	caa Gln ctt Leu atc Ile 5 ttg Leu gaa Glu aat	gca Ala ttt Phe att Ile tct Ser aaa Lys	Met att Ile gac Asp -10 ctt Leu gcc Ala gta Val	116 164 212 260 308 356
ccaa agct cac His gta Val -25 ctg Leu caa Gln atc Ile gat Asp	att Ile -40 cat His gtc Val gaa Glu tat Tyr 25 ctt Leu tgt	tta Leu tgt Cys tgc Cys cag Gln 10 gcc Ala cct Pro	caa Gln cct Pro cat His aaa Lys tca Ser cta Leu	ctg Leu gac Asp gaa Glu -5 aca Thr cag Gln att	ctt Leu act -20 the gtg Val act Thr -20 cta	act Thr -35 gga Gly tgc Cys cta Leu gag Glu agc ser	aca Thr aaa Lys cag Gln gcc Ala 15 caa Gln ctc Leu	gtg Val gac Asp tct Ser tct Ser gag Glu att Ile	gat Asp att Ile gat Asp tat Tyr cgg Arg	gatg atg Asp tTrpsat -15 Asp tte CLeu gtal gag	gga Gly -30 aat Asn cca Pro tca Ser aag Lys 35 tta Leu	att Ile tta Leu ccc Pro gtg Val 20 ata Ile caa Gln aac	caa Gln ctt Leu atc Ile 5 ttg Leu gaa Glu aat Asn	gca Ala ttt Phe att Ile tct Ser aaa Lys Met	Met att Ile gac Asp -10 ctt Leu gcc Ala gta Val gaa Glu 55 gaa	116 164 212 260 308 356
ccaa agct cac His gta Val -25 ctg Leu caa Gln atc Ile gat Asp	cacca gcca att Ile -40 cat His gtc Val gaa Glu tat Tyr	tta Leu tgt Cys tgc Cys cag Gln 10 gcc Ala cct Pro	caa Gln cct Pro cat His aaa Lys tca Ser cta Leu	ctg Leu gac Asp gaa Glu -5 aca Thr cag Gln att Ile aaa Lys	ctt Leu act -20 the gtg Val act Thr -20 cta	act Thr -35 gga Gly tgc Cys cta Leu gag Glu agc ser	aca Thr aaa Lys cag Gln gcc Ala 15 caa Gln ctc Leu	gtg Val gac Asp tct Ser tct Ser gag Glu att Ile	gat Asp att Ile gat Asp tat Val tat Tyr cgg Arg gca Ala	gatg atg Asp tTrpsat -15 Asp tte CLeu gtal gag	gga Gly -30 aat Asn cca Pro tca Ser aag Lys 35 tta Leu	att Ile tta Leu ccc Pro gtg Val 20 ata Ile caa Gln aac	caa Gln ctt Leu atc Ile 5 ttg Leu gaa Glu aat Asn	gca Ala ttt Phe att Ile tct Ser aaa Lys atg Met	Met att Ile gac Asp -10 ctt Leu gcc Ala gta Val gaa Glu 55 gaa	116 164 212 260 308 356 404
ccaa aget cac His gta Val -25 ctg Leu caa Gln atc Ile gat Asp 40 cag Gln act	cacca cacca att le -40 cat His gtc Val gaa tTyr 25 theu tgts aaa	tta Leu tgt Cys tgc Cys cag Gln 10 gcc Ala cct Pro cag	caa Gln cct Pro cat His aaa Lys tca Ser cta Leu aaa Lys	ctg Leu gac Asp gaa Glu -5 aca Thr cag Gln att Ile aaa Lys	ctt Leu actrate Through The Gasp Val actrate Asp 45 ac Pro	act Thr -35 gga Gly tgc Cys cta Leu gGlu agc Ser gag Glu acc	aca Thr aaa Lys cag Gln gcc Ala 15 caa Gln ctc Leu aac Asn	gtg Val gac Asp tct Ser tct gag Glu att tcg gat	gat gat gasp atle gasp tat tat gasp tat tat Tyr carg gala 65 gat	gaty Asp Tribaty Tribaty Tte Cteu Gtal Gaty Sogglu Ctc	gga Gly -30 aat Asn cca Pro tca Ser aag Lys 35 tta Leu tct Ser	att Ile tta Leu ccc Pro gtg Val 20 ata Gln aac Asn	caa Gln ctt Leu atc Ile 5 ttg Leu gaa Glu aat Asn aca Thr	gca Ala ttt Phe att Ile tctr aaa Lys atg Met gag 70 atc	Met atte Ile gac Asp -10 ctt Leu gcc Ala gta Val gaa Glu tta	116 164 212 260 308 356 404
ccaa aget cac His gta Val -25 ctg Leu caa Gln atc Ile gat Asp 40 cag Gln act	att Ile -40 cat His gtc Val gaa Glu tat Tyr 25 ctt Leu tgt	tta Leu tgt Cys tgc Cys cag Gln 10 gcc Ala cct Pro cag	caa Gln cct Pro cat His aaa Lys tca Ser cta Leu aaa Lys	ctg Leu gac Asp gaa Glu -5 aca Thr cag Gln att Ile aaa Lys	ctt Leu actrate Through The gtg Asp Asp Asp Cca Pro	act Thr -35 gga Gly tgc Cys cta Leu gGlu agc Ser gag Glu acc	aca Thr aaa Lys cag Gln gcc Ala 15 caa Gln ctc Leu aac Asn	gtg Val gac Asp tct Ser tctser gag Glu att Ile tcg Ser gat Asp	gat gat gasp atle gasp tat tat gasp tat tat Tyr carg gala 65 gat	gaty Asp Tribaty Tribaty Tte Cteu Gtal Gaty Sogglu Ctc	gga Gly -30 aat Asn cca Pro tca Ser aag Lys 35 tta Leu tct Ser	att Ile tta Leu ccc Pro gtg Val 20 ata Gln aac Asn	caa Gln ctt Leu atc Ile 5 ttg Leu gaa Glu aat Asn aca Thr	gca Ala ttt Phe att Ile tctr aaa Lys atg Met gag 70 atc	Met atte Ile gac Asp -10 ctt Leu gcc Ala gta Val gaa Glu tta	116 164 212 260 308 356 404 452
ccaa agct cac His gta Val -25 ctg Leu caa Gln atc Ile gat Asp 40 cag Gln act Thr	cacca cacca att le -40 cat His gtc Val gaa tTyr 25 theu tgts aaa	tta Leu tgt Cys tgc Cag Gln gca Ala cct Pro cag Arg	caa Gln cct Pro cat His aaa Lys tca Ctau aaa Lys act T75	ctg Leu gac Asp gaa -5 aca Thr cag Gln att Ile aaa Lys 6gat Asp	ctt ag cttu actr-20 che gtgl actr -20 che -20 ch	act Thr -35 gga Cys Cys Leu gglu agglu aggr Thr	aca Thr aaa Lys Cag Gln gcc Ala 15 caa Gln ctc Leu aac Asn cat	gtgl gasp to ser to galu atte to gasp to ser to galu atte to gasp to ser gasp to ser to gasp to gas	gat gasp atle gasp tal tar carg gala try garg cala fasp atle try carg gala fasp at	atg gap gap tribat the cheu cheu cheu cheu cheu cheu cheu ch	gga ggay -30 aatn cca Pro acs acs Leu tcr car his	att Ile tta Leu ccc Pro gtgl 20 ata Ile caa Gln aac Asn ttg Leu cag	caa Gln cttu atce Ile 5ttu gau aat Asn aCar aLys gca	gca Ala ttt Phe att Ectr aaa Lys atg Glu 70 atta	Met atte gac Asp -10 cttu gcc Ala gta Val gaa Glu tta Leu aca	116 164 212 260 308 356 404 452

90 . 95 100	
aag gag acg gtg gct cag gga gta aag gaa ggc cag ttg agc aaa cag	. 596
Lys Glu Thr Val Ala Gln Gly Val Lys Glu Gly Gln Leu Ser Lys Gln 105 110 115	
aag tgt tcc tct gca ttt caa aac ctt ctt cct ttc tat agc cct gtg	644
Lys Cys Ser Ser Ala Phe Gln Asn Leu Leu Pro Phe Tyr Ser Pro Val	
120 125 130 135	
gtg gaa gat ttt att aaa atc cta cgt gaa gtt gat aag gcg ctt gct	692
Val Glu Asp Phe Ile Lys Ile Leu Arg Glu Val Asp Lys Ala Leu Ala	
140 145 150	734
gat gac ttg gaa aaa aac ttc cca agt ttg aag gtt cag act	/34
Asp Asp Leu Glu Lys Asn Phe Pro Ser Leu Lys Val Gln Thr 155 160 165	
taaaacctga attggaatta cttctgtaca agaaataaac tttatttttc tcactgacaa	794
aaaaaaaaaa	804
<210> 89	
<211> 802	
<212> DNA <213> Homo sapiens	
22137 HOMO Bapiens	
<220>	
<221> CDS	
<222> 199801	
<221> polyA_signal	
<222> 780785	
<221> polyA site	
valir polyn_bito	
<222> 791802	
<222> 791802	
<400> 89	.
<400> 89 agtcaccqcc tqcttcqcac tgagcctccc gactcagact ctgagtccag ctccgaagag	60 120
<400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga	120
<400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtgt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctqttatcc qctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc	120 180
<400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtgt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tgtgttggct tgtgttggct tggttggct tggttggct tggttgtgg	120
<400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtgt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctqttatcc qctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc	120 180
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc</pre>	120 180
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala</pre>	120 180 231
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc</pre>	120 180 231 279
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc</pre>	120 180 231
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc</pre>	120 180 231 279
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc</pre>	120 180 231 279
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc</pre>	120 180 231 279
<pre><400> 89 agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc</pre>	120 180 231 279
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 25 caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln 30 35 40 ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile 45 50 55 aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg	120 180 231 279
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 Ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 25 Caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln 30 35 40 Ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile 45 50 55 aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val	120 180 231 279 327
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln 30 35 40 ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile 45 50 55 aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val 60 65 70 75	120 180 231 279 327 375
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln 30 35 40 ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile 45 50 55 aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val 60 65 70 75 aac cac ctc aaa gcc aat gtt aag tca gct gca gac ttg att agc ctg	120 180 231 279 327
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgtggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctgtgtgtggcc tggtgtggct tggtgtgg atg cag gtt gct ctc aag gaa gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 Ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 25 Caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln 30 35 40 Ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile 45 aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val 60 65 70 75 aac cac ctc aaa gcc aat gtt aag tca gct gca gac ttg att agc ctg Asn His Leu Lys Ala Asn Val Lys Ser Ala Ala Asp Leu Ile Ser Leu	120 180 231 279 327 375
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggtgtggct tggtggg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 10 10 10 10 10 10 10 10 10 10 10 10	120 180 231 279 327 375
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggtgtggc atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 25 caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln 30 25 ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile 45 50 55 aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val 60 65 70 75 aac cac ctc aaa gcc aat gtt aag tca gct gca gac ttg att agc ctg Asn His Leu Lys Ala Asn Val Lys Ser Ala Ala Asp Leu Ile Ser Leu B0 cct acc act gta gag gga ctt cag aag agt gta gct tcc att ggc aat	120 180 231 279 327 375 423
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggtgtggct tggtggg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 10 10 10 10 10 10 10 10 10 10 10 10	120 180 231 279 327 375 423
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggttggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggttgtgcct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1 5 10 Ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 25 Caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln 20 25 Ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile 45 50 aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val 60 5 70 75 aac cac ctc aaa gcc aat gtt aag tca gct gca gac ttg att agc ctg Asn His Leu Lys Ala Asn Val Lys Ser Ala Ala Asp Leu Ile Ser Leu 80 90 cct acc act gta gag gga ctt cag aag agt gta gct tcc att ggc aat Pro Thr Thr Val Glu Gly Leu Gln Lys Ser Val Ala Ser Ile Gly Asn 25 act tta aac agc gtc cat ctt gct gtg gaa gct tta aac act gta aac act gta aac act ctt gct gtg gaa gcc tta cag aaa act gtg gaa gcc tta cag aaa act gtg gaa gcc tta cag aaa act gtg gaa gcc ttc att ggc aat pro Thr Thr Val Glu Gly Leu Gln Lys Ser Val Ala Ser Ile Gly Asn 25 act tta aac acc gcc cat ctt gct gtg gaa gcc cta cad aac act gtg	120 180 231 279 327 375 423
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtggt tggaaatcgc tctcgctttg ccaaggggaa ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggtgtggct tggtggg atg cag gtt gct ctc aag gag gat ctg gat gcc kgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc kgtgttggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc kgtgttgtggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc kgtgtgtggct tggtgtgg atg cag gtt gct ctc aag gag gat ctg gat gcc kgtgtgtggct tggtgtggg atg cag atc aac cag aaa agc tca ttc leu lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe log liber libe	120 180 231 279 327 375 423 471
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtgggt tggaaatcgc tctcgctttg ccaagggaga ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggtgttggct tggtgggg atg cag gtt gct ctc aag gag gat ctg gat gcc Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala 1	120 180 231 279 327 375 423 471 519
agtcaccgcc tgcttcgcac tgagcctccc gactcagact ctgagtccag ctccgaagag gaagaggaat tcggtggt tggaaatcgc tctcgctttg ccaaggggaa ctatttacga tgctgcaaga tctgttatcc gctctgtggt tttgtcatcc ttgctgcctg tgttgtggcc tggttggct tggttggg atg cag gtt gct ctc aag gag gat ctg gat gcc gtgtgttggct tggttggg atg cag gtt gct ctc aag gag gat ctg gat gcc ggtggtggtgga atg cag gtt gct ctc aag gag gat ctg gat gcc gtgtgtgtggttggct tggtgtggg atg cag gtt gct ctc aag gag gat ctg gat gcc ggtgtgtggtggtgga atg cag gtt gct ctc aag gag gat ctg gat gcc gtgtgtgtggtggtgga atg cag gtt gat ctg aac aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc leu lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe 15 20 25 25 25 25 25 25 25 25 25 25 25 25 25	120 180 231 279 327 375 423 471

Asp Glu His Lys Lys Thr Met Glu Leu Leu Gln Ser Asp Met Asn Gln	
cac ttc ttg aag gag act cct gga agc aac cag atc att ccg tca cct His Phe Leu Lys Glu Thr Pro Gly Ser Asn Gln Ile Ile Pro Ser Pro 140 145 150 155	663
tca gcc aca tca gaa ctt gac aat aaa acc cac agt gag aat ttg aaa Ser Ala Thr Ser Glu Leu Asp Asn Lys Thr His Ser Glu Asn Leu Lys	711
cag atg ggt gat aga tct gcc act ctg aaa aga cag tct ttg gac caa Gln Met Gly Asp Arg Ser Ala Thr Leu Lys Arg Gln Ser Leu Asp Gln 175 180 185	759
gtc acc aac aga aca gat aca gta aaa atc caa aaa aaa aa aa aa aa aa aa aa aa aa	802
<210> 90 <211> 1490 <212> DNA <213> Homo sapiens	
<220>	
<221> CDS <222> 381174	
<221> sig_peptide <222> 38.148 <223> Von Heijne matrix score 7.3	
seq LLSACLVTLWGLG/EP	
<221> polyA_signal <222> 14521457	
<221> polyA_site <222> 14781490	
<pre><400> 90 tcatcatcca gagcagccag tgtccgggag gcagaag atg ccc cac tcc agc ctg</pre>	55
-35 cat cca tcc atc ccg tgt ccc agg ggt cac ggg gcc cag aag gca gcc His Pro Ser Ile Pro Cys Pro Arg Gly His Gly Ala Gln Lys Ala Ala	103
-30 -25 -20 ttg gtt ctg ctg agt gcc tgc ctg gtg acc ctt tgg ggg cta gga gag Leu Val Leu Leu Ser Ala Cys Leu Val Thr Leu Trp Gly Leu Gly Glu -15 -10 -5 1	151
cca cca gag cac act ctc cgg tac ctg gtc ctc cac cta gcc tcc ctg Pro Pro Glu His Thr Leu Arg Tyr Leu Val Leu His Leu Ala Ser Leu 5 10 15	199
cag ctg gga ctg ctg tta aac ggg gtc tgc agc ctg gct gag gag ctg Gln Leu Gly Leu Leu Asn Gly Val Cys Ser Leu Ala Glu Glu Leu 20 25 30	247
cgc cac atc cac tcc agg tac cgg ggc agc tac tgg agg act gtg cgg Arg His Ile His Ser Arg Tyr Arg Gly Ser Tyr Trp Arg Thr Val Arg 35 40 45	295
gcc tgc ctg ggc tgc ccc ctc cgc cgt ggg gcc ctg ttg ctg tcc Ala Cys Leu Gly Cys Pro Leu Arg Arg Gly Ala Leu Leu Leu Ser	343
50 55 60 65 atc tat ttc tac tac tcc ctc cca aat gcg gtc ggc ccg ccc ttc act Ile Tyr Phe Tyr Tyr Ser Leu Pro Asn Ala Val Gly Pro Pro Phe Thr	391

				70					75					80		
tgg Trp	atg Met	ctt Leu	gcc Ala	ctc	ctg Leu	ggc Gly	ctc Leu	Ser	cag	gca Ala	ctg Leu	aac Asn	atc Ile 95	ctc	ctg Leu	439
ggc Gly	ctc Leu	Lys	85 ggc Gly	ctg Leu	gcc Ala	cca Pro	Ala	90 gag Glu	atc Ile	tct Ser	gca Ala	gtg Val 110	tgt	gaa Glu	aaa Lys	487
gly ggg	aat Asn 115	100 ttc Phe	aac Asn	gtg Val	gcc Ala	cat His	105 999 Gly	ctg Leu	gca Ala	tgg Trp	tca Ser 125	tat	tac Tyr	atc	gga . Gly	535
tat Tyr 130	ctq	cgg Arg	ctg Leu	atc Ile	ctg Leu 135	cca	gag Glu	ctc Leu	cag Gln	gcc Ala 140	cgg	att Ile	cga Arg	act Thr	tac Tyr 145	583
aat	cag Gln	cat His	tac Tyr	aac Asn 150	aac	ctg Leu	cta Leu	cgg Arg	ggt Gly 155	gca Ala	gtg Val	agc Ser	cag Gln	cgg Arg 160	ctg Leu	631
tat Tyr	att Ile	ctc Leu	ctc Leu 165	cca	ttg Leu	gac Asp	tgt Cys	999 Gly 170	gtg Val	cct Pro	gat Asp	aac Asn	ctg Leu 175	agt Ser	atg Met	679
gct Ala	gac Asp	ccc Pro 180	aac Asn	att Ile	cgc Arg	ttc Phe	ctg Leu 185	gat Asp	aaa Lys	ctg Leu	ccc Pro	cag Gln 190	cag Gln	acc Thr	ggt Gly	727
gac Asp	cgt Arg 195	gct	ggc Gly	atc Ile	aag Lys	gat Asp 200	cgg Arg	gtt Val	tac Tyr	agc Ser	aac Asn 205	agc Ser	atc Ile	tat Tyr	gag Glu	775
ctt Leu 210	ctq	gag Glu	aac Asn	GJA aaa	cag Gln 215	cgg Arg	gcg Ala	ggc	acc Thr	tgt Cys 220	gtc Val	ctg Leu	gag Glu	tac Tyr	gcc Ala 225	823
acc	ccc Pro	ttg Leu	cag Gln	act Thr 230	ttg Leu	ttt Phe	gcc Ala	atg Met	tca Ser 235	caa Gln	tac Tyr	agt Ser	caa Gln	gct Ala 240	ggc	871
ttt Phe	agc Ser	cgg Arg	gag Glu 245	gat	agg Arg	ctt Leu	gag Glu	cag Gln 250	gcc Ala	aaa Lys	ctc Leu	ttc Phe	tgc Cys 255	cgg Arg	aca Thr	919
ctt Leu	gag Glu	gac Asp 260	atc Ile	ctg Leu	gca Ala	gat Asp	gcc Ala 265	cct Pro	gag Glu	tct Ser	cag Gln	aac Asn 270	aac Asn	tgc Cys	cgc Arg	967
ctc Leu	att Ile 275	gcc	tac Tyr	cag Gln	gaa Glu	cct Pro 280	gca Ala	gat Asp	gac Asp	agc Ser	agc Ser 285	ttc Phe	tcg Ser	ctg Leu	tcc Ser	1015
cag Gln 290	gag Glu	gtt Val	ctc Leu	cgg Arg	His	Leu	Arg	cag Gln	Glu	Glu	Lys	Glu	gag Glu	gtt Val	acc Thr 305	1063
qtq	qqc	agc Ser	ttg Leu	aag Lys 310	acc	tca	gcg	gtg	ccc	agt	acc	tcc	acg Thr	atg Met 320	tcc Ser	1111
caa Gln	gag Glu	cct Pro	gag Glu 325	ctc Leu	ctc Leu	ctc Leu	agt Ser	gga Gly 330	atg Met	gga Gly	aag Lys	ccc Pro	ctc Leu 335	cct Pro	ctc Leu	1159
			ttc Phe			gacc	cag	ggtc	acca	gg c	caga	gcct	c ca	gtgg	tctc	1214
caa	gcct	ctg	gact	9999	gc t	ctct	tcag	t gg	ctga	atgt	cca	gcag	agc	tatt	tccttc	1274
cac	aggg	ggc	cttg	cagg	ga a	gggt	ccag	g ac	ttga	catc	tta	agat	gcg	tctt	gtcccc	1334
ttg	ggcc	agt	catt	tccc	ct c	tctg	agcc	t cg	gtgt	cttc	aac	ctgt	gaa	atgg	gatcat	1394
aat	cact	gcc	ttac	ctcc	ct c	acgg	ttgt	t gt	gagg	actg	agt	gtgt	gga	agtt	tttcat	1454 1490
aaa	cttt	gga	tgct	agtg	ta c	ссаа	aaaa	a aa	adāā							1430

<212> DNA <213> Homo sapiens	٠.
<220> <221> CDS <222> 26361	
<221> polyA_site	
<pre><400> 91 tcgagaagct gccccttagc caacc atg ccg tct gag ggt cgc tgc tgg gag Met Pro Ser Glu Gly Arg Cys Trp Glu 1</pre>	52
acc ttg aag gcc cta cgc agt tcc gac aaa ggt cgc ctt tgc tac tac Thr Leu Lys Ala Leu Arg Ser Ser Asp Lys Gly Arg Leu Cys Tyr Tyr	100
cgc gac tgg ctg ctg cgc gag gat gtt tta gaa gaa tgt atg tct Arg Asp Trp Leu Leu Arg Arg Glu Asp Val Leu Glu Glu Cys Met Ser	148
ctt ccc aag cta tct tct tat tct gga tgg gtg gta gag cac gtc cta Leu Pro Lys Leu Ser Ser Tyr Ser Gly Trp Val Val Glu His Val Leu 45 50 55	196 244
ccc cat atg cag gag aac caa cct ctg tct gag act tcg cca tcc tct Pro His Met Gln Glu Asn Gln Pro Leu Ser Glu Thr Ser Pro Ser Ser 60 65 70	292
acg tca gct tca gcc cta gat caa ccc tca ttt gtt ccc aaa tct cct Thr Ser Ala Ser Ala Leu Asp Gln Pro Ser Phe Val Pro Lys Ser Pro 80 85	340
gac gca agc tct gcc ttt tcc cca gcc tcc cct gca aca cca aat gga Asp Ala Ser Ser Ala Phe Ser Pro Ala Ser Pro Ala Thr Pro Asn Gly 90 95 100 105	361
acc aag ggc aaa aaa aaa Thr Lys Gly Lys Lys Lys 110	
<210> 92 <211> 605 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 3131	
<221> polyA_site <222> 591605	
<pre><400> 92 ca tcc ctt ccc cag gct tta tgg ttc cag ttc ttc tac cac tct gga Ser Leu Pro Gln Ala Leu Trp Phe Gln Phe Phe Tyr His Ser Gly 1</pre>	47
ago too ota gaa tot oot gga atg ott aat gga oot tto dag dad oga Ser Ser Leu Glu Ser Pro Gly Met Leu Asn Gly Pro Phe Gln His Arg 20 25 30	95
aat toa aga att atg act cat ogg toa goa gaa aag tgaggataco Asn Ser Arg Ile Met Thr His Arg Ser Ala Glu Lys 40	141
ttttcctaac ctacctgctt cccctgcagt ttcctcacaa tcttactctt tatattttag catatgtagc ttctcaggat gttaattctg ttctctctgt gttggtgtct gagcacccag	201 261

aaggtagagc caggggcact tataaaccag gagcattatt tgacaggcac ttaagaaaga cactggctac gtaatcccag cactttggga ggctgaggcg gatggatcac atgaggtcag gagttcgaga ccagcctggc cagcatggtg aaaccctgtc tctactaaaa atacaaaaat tagctgggtg tggttgcaca cgcctgtaat cccagctacc tgggaggctg aggcaggaga atcgcttgaa cttgggaggc ggaggttgca gtgagcctag attttgccat tgcactccag cctgggtgac aagggcgaaa ctccatccca aaaaaaaaaa	321 381 441 501 561 605
<210> 93 <211> 591 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 33185	
<221> sig_peptide <222> 3380 <223> Von Heijne matrix score 3.7 seq IALTLIPSMLSRA/AG	
<221> polyA_signal <222> 570575 <221> polyA_site	
<222> 586591	
<pre><400> 93 caatcttctc agcttataac cgtctttccc tt atg cta agg ata gcc ctt aca</pre>	53
ctc atc cca tct atg ctg tca agg gct gct ggt tgg tgc tgg tac aag Leu Ile Pro Ser Met Leu Ser Arg Ala Ala Gly Trp Cys Trp Tyr Lys	101
gag ccc act cag cag ttt tct tac ctt tgc ctg ccc tgc ctt tca tgg Glu Pro Thr Gln Gln Phe Ser Tyr Leu Cys Leu Pro Cys Leu Ser Trp 15 20	149
aat aag aaa ggc aac gtt ttg cag ctt cca aat ttc tgaagaaact Asn Lys Lys Gly Asn Val Leu Gln Leu Pro Asn Phe	195
25 30 35 aatotoagat tggcagttaa agtoaaaatg ttgccaaata tttattoott ttgcctaagt	255
aatotoagat tggcagttaa agtcaaaatg ttgacaatot gactottoag agttogtaco ttggctacco ggttcaattg ctttttattt ttaatgtott gactottoag agttogtacc	315
	375
	435
	495 555
tgccaggagc tccttccctc tagcaattee tuctaatateg agtaaatcat tttggtagct ttacaatcaa attactgtat ttattaattt gctagaatcc agtaaatcat tttggtagct ctggctgtgc tatcaataaa aagatgaaag caaaaa	591

<210> 94 <211> 1150 <212> DNA <213> Homo sapiens

<220>

<221> CDS <222> 184..915

<221>				₹												
<222				ma+1	ri v									•		••
<223		ore:		macı	LIX											
	sec	JLE .	3.5 GLEL	SEAE	AIG/	AD										
<221:		_														
<222	_	_								•						
<221: <222:																
<400	> 94								- 1 -				a-	+ < + +	acatt	60
cgga	tttg	ac g	atgg	tgtt	c gg	tctt	gaat	gga	aatg	tag	2020	agge	ca y	taga	aggtt	-
tttg	aaca	gg a	tagt	aggt	a tc	cgga	gtcg	act +c+	gagg	gee	agag	agat	ct c	tttc	gttcg	
gate gcc	ctgg	gc a	aagt	ttcc	c ac	gttg	ayyy cta	aga	cta	gac	cta	agc	gag	gcg	ccagg gag	228
gcc	atg '	gcg (aac Aan	Dro '	aay Lvs	Leu	Leu	Gly	Leu	Glu	Leu	Ser	Glu	Ala	Glu	
				-15					-10					-5		
gcg	atc	ggt	gct	gat	tcg	gcg	cga	ttt	gag	gag	ctg	ctg	ctg	cag	gcc	276
Ala	Ile	Gly	Ala 1	Asp	Ser	Ala	Arg 5	Phe	Glu	Glu	Leu	10	ьеи	GIII	AIG	204
tcg	aag	gag	ctc	cag	caa	gcc	cag	aca	acc	aga	cca	gaa	tcg	aca	caa	324
Ser	Lys 15	Glu	Leu	Gln	Gln	Ala 20	Gln	Thr	Thr	Arg	Pro 25	Glu	Ser	Thr	GIN	
2+0	cac	cct	cag	cct	aat	ttc	tgc	ata	aag	acc	aac	tcc	tcg	gaa	9 99	372
Ile	Gln	Pro	Gln	Pro	Gly	Phe	Cys	Ile	Lys	Thr	Asn	Ser	Ser	Glu	Giy	
3.0					3.5					40					43	420
aag	gtt	ttc	atc	aac	atc	tgc	cac	tcc	CCC	tct	atc	cct	cct	CCC	gcc	420
				50					55			Pro		60		
qac	gtg	acc	gag	gag	gag	ctg	ctt	cag	atg	cta	gag	gag	gac	caa	gct	468
Asp	Val	Thr	Glu 65	Glu	Glu	Leu	Leu	Gln 70	Met	Leu	Glu	Glu	Asp 75	Gln	Ala	
aaa	+++	cac	atc	ccc	atq	aqt	ctg	gga	gag	cct	cat	gca	gaa	ctg	gat	516
Gly	Phe	Arg	Ile	Pro	Met	Ser	Leu	Gly	Glu	Pro	His	Ala 90	Glu	Leu	Asp	
		80					85	+=0	cac	ata	act		aac	agc	gac	564
gca	aaa	ggc	cag	gga	Cyc	Thr	Δla	Tvr	Asp	Val	Ala	gtc Val	Asn	Ser	Asp	
Ата	шуs 95	GIY	GIII	Gry	Cys	100	ALU	-1-			105					
ttc	tac	caa	agg	atq	caq	aac	agc	gat	ttc	ttg	cgg	gag	ctc	gtg	atc	612
Phe	Tvr	Arg	Arg	Met	Gln	Asn	Ser	Asp	Phe	Leu	Arg	Glu	Leu	Val	TIE	
110					115					120					125	660
acc	atc	gcc	agg	gag	ggc	ctt	gag	gac	ata	tac	aac	ttg	cag	ctg	aat	660
Thr	Ile	Ala	Arg	Glu	Gly	Leu	Glu	Asp	TIE	Tyr	ASD	Leu	GIII	140	ASII	
				130			224	caa	135	ttc	ato	ggc	taa		tca	708
ccg	gaa	tgg	cgc	atg	Met	Lve	Agn	Arg	Pro	Phe	Met	ggc Gly	Ser	Ile	Ser	
			145					150					155			
cag	cag	aac	atc	cgc	tcg	gag	cag	cgt	cct	cgg	atc	cag	gag	ctg	a aa	756
Gln	Gln	Asn	Ile	Arg	Ser	Glu	Gln 165	Arg	Pro	Arg	IIe	170	GIU	ъеп	GIY	
gac	cta	tac	acg	ccc	gcc	ccc	999	aga	gct	gag	tca	999	cct	gaa	aag	804
Asp	Leu 175	Tyr	Thr	Pro	Ala	Pro 180	Gly	Arg	Ala	Glu	Ser 185	GIY	Pro	GIU	пув	
cct	cac	cta	aac	cta	tga	ctq	gaa	gcc	ccc	gac	ctc	ctc	ttg	gcc	gaa	852
Pro	His	Leu	Asn	Leu	Trp	Leu	Glu	Ala	Pro	Asp	Leu	Leu	Leu	Ala	Giu	
190					195					200					205	900
gtt	gac	ctc	CCC	aaa	ctg	gat	gga	gco	ctg	999	ctg	tcg	ctg	gag	atc	300
Val	Asp	Leu	Pro			Asp	GIÀ	Ala	Leu 215	: r GTÅ	ьeu	. ser	TEO	220	Ile	
~~~	2~-	200		210	tas	taaa	aaa	cccc			tata	tcat	c ta		ctta	955
999	aya	acc	gee	-33	-ya	-222	223		5-	-5 -	- 5					

Gly Ar		2	っち													٠.
tatcco attaat tctgag	gatg	g ca g gc t gg	gato catg agaa	CCCC	ttc	tacc	aat	acct	tctt	ga t	cagg	gugu	C CC	cccg	,-5	1015 1075 1135 1150
tttaaa	aaaa	a aa	aaa													2204
<210>																
<211>																
<212><213>			nier	ıs												
<220>		.0 30	.p.c.													
<221>		}														
<222>			16													
<221>				3												
<223>				mat	rix											
		ore 4			·····	20										
				HLYD	VFG/1	JP										
<221>	po]	lyA_: 86	sign 1491	al												
<221>	. ກດ ⁻	4v1	site													
<222>	_	_														
<400: ctga	+	ta a	atto	tcac	a ac	actt	gacc	aat	aaga	ttc	ggga	gctt	ct t	cagc	aa	57
	~	202	~~~	cta	222	tca	aca	qac	cct	cgg	gat	ggc	acc	990	cuo	105
Met (	3lu 1	Arg	Gly	Leu	Lys	Ser	Ala	Asp	Pro -25	Arg	Asp	GIY	TILL	-20	- 7 -	153
act 9	ggc	tgg	gca	ggt	att	gct	gtg	ctt	tac	tta	cat	ctt	tat	gat	gta Val	100
Thr (	Gly '	Trp	Ala	Gly	Ile	Ala	Val	Leu -10	Tyr	Leu	HIS	пец	-5	No P	· · · ·	
ttt	700	a a c	-15	acc	tac	cta	caq	tta	gca	cat	ggc	tat	gta	aag	caa	201
Phe	gly .	Asp	Pro	Ala	Tyr	Leu	Gln	Leu	Ala	His	GIY	Tyr	Val	Lys	Gln	
		9				5					TU					249
agt Ser	ctg	aac	tgc	tta	acc	aag	cgc	tcc	atc	Thr	Dhe	Leu	Cvs	Glv	Asp	
	Leu	Asn	Cys	Leu	20	пуs	Arg	361	110	25			-2-	•	30	
15 gca	aac	ccc	cta	qca	ata	qcc	gct	gtg	cta	tat	cat	aag	atg	aac	aat	297
Ala	Gly	Pro	Leu	Ala	Val	Āla	Ala	Val	Leu	Tyr	His	Lys	Met	Asn 45	Asn	
gag			~~~	35	ant.	tac	atc	aca	40 cga	cta	att	cac	cta		aag	345
gag Glu	aag Lys	Gln	Ala	Glu	Asp	Cys	Ile	Thr	Arg	Leu	lle	His	Leu 60	Asn	Lys	
a++	ant.	cct	50	act	сса	aat	gaa	atq	ctc	tat	999	cga	ata	ggc	tac	393
Ile	Asp	Pro	His	Ala	Pro	Asn	Glu 70	Met	Leu	Tyr	GIY	75	116	GIY	-y-	
atc	tat		ctt	ctt	ttt	gtc	aat	aag	aac	ttt	gga	gtg	gaa	aag	act	441
Ile	Tyr	Ala	Leu	Leu	Phe	Val 85	Asn	Lys	Asn	Pne	90	vai	GIU	пув	1111	400
cct		agc	cat	att	cag	cag	att	tgt	gaa	aca	att	tta	acc	tct	gga	489
Pro	Gln	Ser	His	Ile	Gln	Gln	Ile	Cys	Glu	Thr 105	TIE	ren	Tur	ser	Gly 110	
95				2~~	100	202	220	ttc	acc			tct	cca	ctg	atg	537
gaa Glu	aac Asn	Leu	Ala	agg Arg 115	Lys	Arg	Asn	Phe	Thr 120	Ala	Lys	Ser	Pro	Leu 125	1100	

		4	tac		~	+=+	tat	ata	aaa	act	act	cat	aac	ctg	gct	585
tat	gaa	tgg	Tyr	cay	Clu	Tur	Tur	Val	GJA	Ala	Ala	His	Gly	Leu	Ala	
Tyr	GIU	Trp		GIII	Gru	TYL	-1-	135	<b>-</b> 2				140			
			130 tac	+ > 0	cta	ato	cag	CCC	agc	ctt	caa	qtq	agc	caa	ggg	633
gga	att	tat	Tyr	Tur	Len	Met	Gln	Pro	Ser	Leu	Gln	Val	Ser	Gln	Gly	
GIÀ	TIE	Tyr	TYL	IÀT	пец	Mec	150					155				
		145	agt		ata	220	220	agt	αt.a	gac	tac	atc	tgc	cag	ctg	681
aag	tta	cat	Ser	Tou	wal	Luc	Dro	Ser	Val	Asp	Tvr	Val	Cys	Gln	Leu	
Lys		HIS	Ser	пеп	vai	165	FIO	001			170		•			
	160		tct	~~~	22 t	tac	cct	cca	tat	ata	aat	gat	aat	cga	gat	729
aaa	בנכ	200	Ser	990	Agn	Tyr	Pro	Pro	Cvs	Ile	Glv	Asp	Asn	Arg	Asp	
	Pne	Pro	Ser	GIY	180	- y -	110		0,70	185		-			190	
175			cat	+ aa	+00	cat	aac	acc	cct		qta	atc	tac	atg	ctc	777
ctg	CEE	gcc	His	~~~	Cyc	Hie	Glv	Ala	Pro	Glv	Val	Ile	Tyr	Met	Leu	
Leu	ьeu	vaı	HIS	195	Cys	1113	<b>G 1 1</b>		200	2			•	205		
			tat	193	at a	ttc	aga	gag		aaq	tat	ctc	tgt	gat	gcc	825
atc	cag	310	Tyr	Tys	Val	Dhe	Ara	Glu	Glu	Lvs	Tyr	Leu	Cys	Asp	Ala	
TIE	GIN	Ald	210	пуъ	Val	1110	••••	215			•		220			
		+~+	210	gat	ata	atc	taa	caa	tat	qqq	ttg	ctg	aag	aag	gga Gly	873
tat	cag	Cyc	712	Acn	Val	Tle	Trp	Gln	Tyr	Gly	Leu	Leu	Lys	Lys	Gly	
TYE	GIN	225	MIA	Asp	٧۵٢	110	230		- 2 -	•		235				
	~~~		tac	cac	aat	tet	gca	aaa	aat	qcc	tat	gcc	ttc	ctg	aca Thr	921
mir	999	Len	Cyc	His	Glv	Ser	Ala	Gly	Asn	Ala	Tyr	Ala	Phe	Leu	Thr	
	240					245					250					
ot c	+ 20	220	ctc	aca	cag	gac	atq	aag	tac	ctg	tat	agg	gcc	tgt	aag	969
Ton	Tur	Agn	Len	Thr	Gln	Asp	Met	Lys	Tyr	Leu	Tyr	Arg	Ala	Cys	Lys 270	
255					260					265					270	
+++	act	gaa	t.aa	tac	tta	qaq	tat	gga	gaa	cat	gga	tgc	aga	aca	cca	1017
Dhe	Δla	Glu	מיד ו	Cvs	Leu	Glu	Tyr	Gly	Glu	His	Gly	Cys	Arg	Thr	Pro	
				275					280					205		
gac	acc	cat	tto	tct	ctc	ttt	gaa	gga	atg	gct	999	aca	ata	tat	ttc Dhe	1065
Asn	Thr	Pro	Phe	Ser	Leu	Phe	Glu	Gly	Met	Ala	. Gly	Thr	Ile	Tyr	Phe	
			290)				295	,				300	,		
cta	act	gac	cto	cta	gto	ccc	aca	aaa	gcc	agg	tto	cct	gca	ttt	gaa Glu	1113
Leu	Ala	Ast	Leu	Leu	ı Val	Pro	Thr	Lys	. Ala	Arg	Phe	יום פ	, MT	. Phe	Glu	
		305	;				310)				315)			
cto	: t.ga	aago	gata	gcat	gcca	icc t	gcaa	ictca	ac to	cate	gacco	ttt	ctgt	ata		1166
T 033																
++-		cca	agct	aagt	gc t	tccg	ttgo	t tt	ccaa	ıggaa	aca	aaaga	igtc	aaac	tgtgga	1226
	+	++0	ttac	*~++	-++ t	caga	attt	a to	ctttc	catto	agu			Cari	accure	1286
+ = 0			taga	aata	atc c	caago	raaqt	c tt	ttaa	ictti	: aat		det		, CC Caaa	1346
~~~		*+~=	ata:	+ = + c	rta c	ragto	rttt	a ac	atto	qtata	a cat	atat		ayac	accegga	1406
aas	aato	tta	ttta	aaqtt	ta t	gaat	ataa	ac ca	atoto	gttac	tgi		iaaa	atgt	ttaaaa	1466
gaa	acto	caat	acag	gataa	aag a	ataaa	atato	gt ga	actat	taaa	a aaa	aaaaa	ì			1513
			•	-	-											

<210> 96

<211> 417

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 327..416

<221> polyA_site <222> 404..417

<400> 96

tgttttgagg tgttggcatt cttcgctgat ttggctgttc ccaatgttta cattatttaa 60 tottgcaaaa atggttotgt gcacttggat gtgaaatgct gtccagtttt attttttta 120

tgttg atttt aggaa ttgct	ggtc tgat gcaa	a to a ca g ct	ttta taga atct	igaag iaaag :gccc	tta att aag	tcag ttat jtta	gaa ttt atg Met	aaaa caa Gln	tgag atg Met	gtt g gac Asp	taa aca Thr	agctt ttt Phe	g to ttt Phe	atg Met	tca Ser	240 300 353
gaa a Glu L	aa c ys H	ac a	ca c	lis T	ca c hr E	ac a His T	aca o	at a	rie i	cac a His :	rhr l	cac a His '	rhr I	7+9 ·	aaa Lys 25	401
10 aca a Thr L			ys I	aaa a												417
<210>		1														
<2112																
<213	> Hon	no s	apie	ns												
<220		-														
<2213			8													
<221: <222: <223:	> 63 > Vo	20	6 ijne		rix											
				LLFG	SLA/	GL										
	cctt	30 0	20 2	ccgg ct g	ac t	ca o	rta o	ita c	ct t	tq c	at i	-99 4	he G	ly F	agagaa :tt Phe	60 107
			_	45				-	40				-	. 3 3		155
ggc Gly	Tyr	Ala	Ala	Leu	Val	Ala	Ser	Gly -25	Gly	IIe	TIE	GIY	-20	Val	пуз	
Ala	Gly	Ser	gtg Val	ccg Pro	Ser	Leu	Ala -10	Ala	GIĀ	ьeu	ьeu	-5	GIY	361	БСС	203
gcc Ala	Gly	ctg Leu	ggt Gly	gct Ala	tac Tyr 5	cag Gln	ctg Leu	tct Ser	cag Gln	gat Asp 10	cca Pro	agg Arg	aac Asn	gtt Val	tgg Trp 15	251
gtt Val	l ttc Phe	cta Leu	gct Ala	aca Thr	tct	ggt Gly	acc Thr	ttg Leu	Ala	ggc	att Ile	atg Met	gga Gly	atg Met 30	agg Arg	299
++0	tac	C2C	tct	20 gga	222	ttc	ato	cct	25 qca	qqt	tta	att	gca	ggt	gcc	347
Phe	Tyr	His	Ser	Gly	Lys	Phe	Met	Pro	Ala	Gly	Leu	Ile	Ala 45	Gly	Ala	
aqt	ttg	ctg	ato	gtc	gcc	aaa	gtt	qqa	gtt	agt	atg	ttc	aac	aga	ccc	395
Ser	Leu	Leu	Met	Val	Ala	Lys	Val 55	Gly	Val	Ser	Met	Phe 60	Asn	Arg	Pro	
cat	tage	50 caga:	agt	catg	ttcc	ag c		actg	a tg	aaga	atta		atct	gca		448
Hig															acattt	508
teti	tcca:	cta :	tttt aaaa	caat: gaca	at a cc a	ctaa aact	yaga tggc	a ga	gagg	tgga	aaa	tcag	tca	tgat	tacaaa	561
ccta	acag	agg	tggc	gagt	at g	taac	acaa	g ag	ctt							60:

<211> 522 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 2163	
<221> polyA_signal	
<221> polyA_site <222> 511522	
<pre>&lt;400&gt; 98 c gag att gcg ggc tat ggc gcc gaa ggt ttt tcg tca gta ctg gga tat Glu Ile Ala Gly Tyr Gly Ala Glu Gly Phe Ser Ser Val Leu Gly Tyr 1</pre>	49
ccc cga tgg cac cga ttg cca ccg caa agc cta cag cac cac cag tat Pro Arg Trp His Arg Leu Pro Pro Gln Ser Leu Gln His His Gln Tyr 20 25 30	97
tgc cag cgt cgc tgg cct gac cgc cgc tgc cta cag agt cac act caa Cys Gln Arg Arg Trp Pro Asp Arg Arg Cys Leu Gln Ser His Thr Gln	145
tcc tcc ggg cac ctt cct nntgaaggag tggctaaggt tggacaatac Ser Ser Gly His Leu Pro	193
acgttcactg cagctgctgt cggggccgtg tttggcctca ccacctgcat cagcgcccat gtccgcgaga agcccgacga ccccctgaac tacttccccg gtggctgcgc cnggaggcct gactctggga gcacgcacgc acaactacgg gattggcgcc gccgcctgcg tgtactttgg catagcggcc tccctggtca agatgggccg gctggagggc tgggaggtgt ttgcaaaacc caaggtgtga gccctgtgcc tgccgggacc tccagcctgc agaatgcgtc cagaaataaa ttctgtgtct gtgtgtgaaa aaaaaaaaa	253 313 373 433 493 522
<210> 99 <211> 956 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 13465	
<221> sig_peptide <222> 1375	
<pre>&lt;400&gt; 99 ngagtcggga aa atg gct gcg agt acn tcn atg gnc ccg gtg gct gtg acg</pre>	51
gcg gca gtg gcg cct gtc ctg tcc ata aac agc gat ttc tca gat ttg Ala Ala Val Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu -5	99
cgg gaa att aaa aag caa ctg ctg ctt att gcg ggc ctt acc cgg gag Arg Glu Ile Lys Lys Gln Leu Leu Leu Ile Ala Gly Leu Thr Arg Glu 10 15 20	147
egg gge eta eta eac agt age aaa tgg teg geg gag ttg get tte tet	195

• •	
Arg Gly Leu Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser	
25 30 35 40 ctc cct gca ttg cct cnt ggc cag ctg caa ccg cct ccg cct att aca	243
ctc cct gca ttg cct cnt ggc cag ttg cal tcg cc ccg bro Pro Pro Pro Pro Pro Pro Pro Pro Pro P	
Leu Pro Ala Leu Pro Xaa Gly Gln Leu Gln Pro Pro Pro Pro Ile Thr	
45	291
gag gaa gat gcc cag gat atg gat gcc tat acc ctg gcc aag gcc tac	2,7
Glu Glu Asp Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr	
60 65 70	220
ttt gac gtt aaa gag tat gat cgg gca gca cat ttc ctg cat ggc tgc	339
Phe Asp Val Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys	
75 80 85	205
aat agc aag aaa gcc tat ttt ctg tat atg tat tcc aga tat ctg gtg	387
Asn Ser Lys Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val	
90 95 100	
agg gcc att tta aaa tgt cat tct gcc ttt agt gaa aca tcc ata ttt	435
Arg Ala Ile Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe	
105 110 115 120	
aga acc aat gga aaa gtt aaa tct ttt aaa tagcttagca gtgggccact	485
Arg Thr Asn Gly Lys Val Lys Ser Phe Lys	
125 130	
gaatgaatgt actttataca tagcaataat aaaaaaaaga tatcataaat aaagttaaaa	545
aggatggtag agaagaaaat attottagga atgactaaca ggataagtaa caacotgatt	605
atttatttac tttaggttat ataaggttct tcatgcctgt gaattaatat tattgtgtaa	665
gaattaagtt aaaaagcctg ggctgacttt taaatttata aattcattta tcatgtttat	725
agtatattta ttgtttttct ttcatggcta ttaaaaagta tgactgtaaa ggacaatgca	785
agnaaaccaa cttaatactg tattgaataa taagtacaat ttattatttt actttgaaac	845
attatgaatt tactttccta ctttttctta gttgttatct atataaattg attaaaaaaa	905
cattttatgt acntnncatt tectagtaca ggttgagtat ceettatttg a	956
Cattleage achimicall tectageaca ggeogageac coordinates	
<210> 100	
<211> 1041	
<212> DNA	-
<213> Homo sapiens	
(215) None Dapton	
<220>	
<221> CDS	
<222> 20703	
(2227 20/03	
221. sie nestide	
<221> sig_peptide	
<222> 2094	
<223> Von Heijne matrix	
score 3.9	
seq ATVGLLMLGVTLP/NS	
<221> polyA_signal	
<222> 10001005	
<221> polyA_site	
<222> 10231041	
<400> 100	
cagggtectg catectace atg teg atg get gtg gaa ace ttt gge tte tte	52
Met Ser Met Ala Val Glu Thr Phe Gly Phe Phe	
-25 -20 -15	
atg gca act gtg ggg ctg ctg atg ctg ggg gtg act ctg cca aac agc	100
Met Ala Thr Val Gly Leu Leu Met Leu Gly Val Thr Leu Pro Asn Ser	
-10 -5	
tac tgg cga gtg tcc act gtg cac ggg aac gtc atc acc acc acc	148
Tyr Trp Arg Val Ser Thr Val His Gly Asn Val Ile Thr Thr Asn Thr	
5 10 15	

atc Ile	ttc Phe 20	gag Glu	aac Asn	ctc Leu	tgg Trp	ttt Phe 25	agc Ser	tgt Cys	gcc Ala	acc Thr	gac Asp 30	tcc Ser	ctg Leu	ggc Gly	gtc Val	196
tac Tyr 35	aac	tgc Cys	tgg Trp	gag Glu	ttc Phe 40	ccq	tcc Ser	atg Met	ctg Leu	gcc Ala 45	ctc	tct Ser	gly ggg	tat Tyr	att Ile 50	244
cag Gln	Ala	Cys	Arg	Ala 55	ctc Leu	Met	Ile	Thr	Ala 60	Ile	Leu	Leu	Gly	ttc Phe 65	Leu	292
Gly	Leu	Leu	Leu 70	Gly	Ile	Ala	Gly	Leu 75	Arg	Cys	Thr	Asn	11e 80	gly ggg	GIY	340
Leu	Glu	Leu 85	Ser	Arg	Lys	Ala	Lys 90	Leu	Ala	Ala	Thr	Ala 95	Gly	gcc Ala	Pro	388
cac His	att Ile 100	ctg Leu	gcc Ala	ggt Gly	atc Ile	tgc Cys 105	Gly 393	atg Met	gtg Val	gcc Ala	atc Ile 110	tcc Ser	tgg Trp	tac Tyr	gcc Ala	436
ttc Phe 115	aac	atc Ile	acc Thr	cgg Arg	gac Asp 120	ttc Phe	ttc Phe	gac Asp	ccc Pro	ttg Leu 125	tac Tyr	ccc Pro	gga Gly	acc Thr	aag Lys 130	484
tac	gag Glu	ctg Leu	ggc Gly	ccc Pro 135	gcc	ctc Leu	tac Tyr	ctg Leu	ggg Gly 140	tgg Trp	agc Ser	gcc Ala	tca Ser	ctg Leu 145	atc Ile	532
tcc Ser	atc Ile	ctg Leu	ggt Gly 150	qqc	ctc Leu	tgc Cys	ctc Leu	tgc Cys 155	tcc Ser	gcc Ala	tgc Cys	tgc Cys	tgc Cys 160	ggc	tct Ser	580
gac Asp	gag Glu	gac Asp 165	cca Pro	gcc Ala	gcc Ala	agc Ser	gcc Ala 170	cgg Arg	cgg Arg	ccc Pro	tac Tyr	cag Gln 175	gct Ala	cca Pro	gtg Val	628
tcc Ser	gtg Val 180	atg Met	ccc	gtc Val	gcc Ala	acc Thr 185	tcg Ser	gac Asp	caa Gln	gaa Glu	ggc Gly 190	gac Asp	agc Ser	agc Ser	ttt Phe	676
ggc Gly 195	aaa Lys	tac	gly	aga Arg	aac Asn 200	gcc Ala	tac Tyr	gtg Val	tag	cagc		ggcc	cgtg	gg		723
								. ~~	~~~~	a+ aa	~~~	aaac	cca	++cc	cctata	783
CCC	cgct	gtc	ttcc	cact	gc c	ccaa	9949	a 99	gyac	229	acc	2226	cca	caac	cctata	843
gta	acct	cag	gggc	cggc	ca c	gccc	cgct	C CC	gtag	4500	900	~==~	tca	cate	cccgtg	903
tct	tgca	ctc	tcat	ggcc	cc t	ccag	gcca	a ga	actg		- tgg	yaag tata	220	caca aaaa	tetece	963
ctc	tgag	gct	ggat	ccct	ca t	CTTC	rgac	C CE	9995	actg	990	cgcg	aay =++	atas	cggtgt ccgtta	1023
		.cgt .aaa				ataa	atac	a tt	Catt	aalā	aat	gcac	400	grya	ccgtta	1041

<210> 101 <211> 558 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 103..294 <221> sig_peptide <222> 103..243 <223> Von Heijne matrix

score 5.9

seq TWLGLLSFQNLHC/FP

<400> 101

WO 99/31236	,	-77-	PCT/IB98/02122
taacattaac ttcctt	.aagt aataatcaat	gaaagaaatt ct at	g cat ggt ttt 114 t His Gly Phe -45
gaa ata ata tcc t Glu Ile Ile Ser I	eu Lys Glu Glu	Ser Pro Leu Gry	lag gtg agt cag 162 Lys Val Ser Gln -30
ggt cct ttg ttt a	aat gtg act agt Asn Val Thr Ser	Gly Ser Ser Ser	ca gtg acc tgg 210
Leu Gly Leu Leu S	-20 ccc ttc cag aac Ser Phe Gln Asn	ctg cat tgc ttc ( Leu His Cys Phe	ca gac ctc ccc 258
Thr Glu Met Pro	cta aga gcc aaa Leu Arg Ala Lys	gga gtc aac act	gagcctagg 304
gtgggctaca acaaa cttgctgaag gaact	taaaa agtagctgtt	ttgcttcatc tagg t atttattgta ttgt	tccagg ccccaagtag 364 ataagc taaaaacatt 424 tgttca cggtgtttgt 484 ggaatt gaccggatag 544 558
<210> 102 <211> 730 <212> DNA <213> Homo sapie	ns		
<220> <221> CDS <222> 81518		÷	
<221> sig_peptid <222> 81173 <223> Von Heijne score 3.9 seq ILFHGV			
<400> 102 ctcgtcatgc tcttt attttcaaga gagtt	gtgct atg atg t Met Met T	Trp Gln Lys Tyr A	acaactt gcotttgatg 60 ca gga agc agg cgg 113 la Gly Ser Arg Arg
tca atg cct ctg Ser Met Pro Leu	Gly Ala Arg 116	c ctt ttc cac ggt Leu Phe His Gly	gtg ttc tat gcc 161
-20 ggg ggc ttt gcc Gly Gly Phe Ala	-15 att gtg tat tac Ile Val Tyr Tyr	-10 c ctc att caa aag r Leu Ile Gln Lys	ttt cat tcc agg 209 Phe His Ser Arg
cat tto tot toc	l laggetta aca ata	5 g gag cag ctg cag l Glu Gln Leu Gln	agc cat ccc gag 257 Ser His Pro Glu
15	20 cta age cet cet	t ctc aac atc cat o Leu Asn Ile His	tat ctc aag ctc 305
30	35	c att gtt gat gcc p Ile Val Asp Ala	aag ttg aag att 353 Lys Leu Lys Ile
45	50 . too aaa toa ga	g ggc ctt ctc tac u Gly Leu Leu Tyr	gtc cac tca tcc 401 Val His Ser Ser
	65	70 g cac ctt gac gag	, ,

Arg Gly Gly Pro Phe Gln Arg Trp His Leu Asp Glu Val Phe Leu Glu 85 90	
ctc aag gat ggt cag cag att cct gtg ttc aag ctc agt ggg gaa aac Leu Lys Asp Gly Gln Gln Ile Pro Val Phe Lys Leu Ser Gly Glu Asn	497
ggt gat gaa gtg aaa aag gag tagagacgac ccagaagacc cagcttgctt Gly Asp Glu Val Lys Lys Glu	548
ctagtccatc cttccctcat ctctaccata tggccactgg ggtggtggcc catctcagtg acagacactc ctgcaaccca gttttccagc caccagtggg atgatggtat gtgccagcac atggtaatt tggtgtaatt ctaacttggg cacaacgaat gctatttgtc attttaaac	608 668 728 730
tg	
<210> 103 <211> 1098	
<212> DNA <213> Homo sapiens	
<220> <221> CDS	
<222> 66326 <221> polyA_signal	
<222> 10661071	
<221> polyA_site <222> 10871098	
<400> 103	60
ctccctttga atgagagaaa ctaacccgct tccgaagccc ctgaaagaca ctgctccttc	
ctctc atg gag ttg gct ccg aca gcc cgt ctg cca cca ggc cat ggt tcc  Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly Ser	110
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly Ala Gly Sch 1 5 10 15 ttg ccc sat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac	110
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly Sci 1 5 10 15 ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His	
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly 3c1  1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20 25 30  etc tct ctt ctc ccc rag atc aag caa cgt gcc tca gag gct ttg ccc	
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly 3c1  1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20 25 30  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro	158
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly 3c1  1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac  Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20 25 30  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc  Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  35 40 45  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln	158
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly  1	158 206
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly 3c1  1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac  Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20 25 30  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc  Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  35 40 45  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag  Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln  50 55 60  tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag  Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu  65 70 75	158 206 254
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly 3ct  1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac  Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20 25 30  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc  Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  35 40 45  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag  Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln  50 55 60  tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag  Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu  65 70 75  ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc geattctcca  Leu Glu Val Asp Asp Trp Glu Phe	158 206 254 302 356
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly 3ct 1 1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His 20 25 30  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro 35 40 45  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln 50 55 60  tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu G5 70 75  ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc gcattctcca Leu Glu Val Asp Asp Trp Glu Phe 80 85	158 206 254 302
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly Als Gly Str  1	158 206 254 302 356 416 476 536
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly Als Gly Str  1	158 206 254 302 356 416 476 536 596
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly Als Gly 15 de 1 1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His 20 25 30   ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro 35 40 45   gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln 50 55 60   tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu Glu 65 70 75   ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc geattctca Leu Glu Val Asp Asp Trp Glu Phe 80 85  gccagggatg cagaggccac ccagaggcc ttcctgaggg ccggccacat tcccgcctc ctgggcagat tgggtagaaa ggacattctt ccaggaaagt tggacacaa gggaatctca acaaccaggg atcaggaggg tccaaagccg acattcccag tcctgtgagc tcagggggt tccaaagccg acattcccag tcctgtgagc tccagggttt tccaaggagtt cccaaggagtt cccaaggagt cccaaggagt cccaaggagt cccaaggagt cccaaggagt cccaaggagt cccaaggagt tccaaagccg acattcccag tcctgtgagc tccaaggatt tccaaggagt cccaaggagt cccaaggagt cccaaggagt cccaaggagt cccaaggagt tccaaggagt tccaaggagt cccaaggagt tccaaggagt tccaaggagt tccaaggagt cccaaggagt c	158 206 254 302 356 416 476 536
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly Als Gly Str 1	158 206 254 302 356 416 476 536 596 656 716 776
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly Als Strong 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac  Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20  25  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc  Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  35  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag  Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln  50  tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag  Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu  65  ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc geattctca  Leu Glu Val Asp Asp Trp Glu Phe  80  gccagggatg cagaggccac ccagaggccc ttcctgagg ccggccacat tcccgcctc  ctgggcagat tgggagata cttcactgct ccaaagcttt tgagacacaa gggaatctca  acaaccaggg atcaggagga tccaaagccg acattcccag tcctggagat tgggagatctca  acaaccaggg atcaggagga ccggcacct tcccaggagct tgactgctgg ctcattctaa  acaaccaggg atcaggagga ccgcacctct tcccaggagct tgactgcagc cccagggttt  ggctccttaa acccgaggac ccgcacctt tcccaggagct tgcaccagc ctcattctac  ttaactttgc tctcagatgc ctccagatgct ataggtcagt gaaagggca gtagtaagct  ggctcgctcc cttccgcag accttctccct cataattcca gagaaggca ttctctcct  cataattcc gagaaggca cttcctccct cataattcca gagaaggca ttctctctct	158 206 254 302 356 416 476 536 596 656 716 776 836
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20 25 30  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  35 40  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln  50 55  tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu  65 70  ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc gcattctca Leu Glu Val Asp Asp Trp Glu Phe  80 85  gccagggatg cagaggaca ccagaggcc ttcctgagg ccggcacat tcccgcctc ctgggcagat tgggtagaaa ggacattctt ccaggaaagt tgactgctgg tcaaagccg acaaccagg atcaggaggt ccaaagccg acattccag cctgggagct tgagacacaa gggaatctca acaaccagg atcaggaggt cccaaagccg acattccag cctgggagct tgagagacacca tcctcccgcag aagagagatg ctgcctctgc cctgggagct tgccaaagccg ctctctcagagg cccacactct tcccaggaact tgagaccaca gggaatctca tcacttgc tcccagatgc ctccagatgct tcccaggacg tcctgggagct gaattccaag gcctccttaa acccgaggac ctccagatgct tcccaggacgt tggaccagc ctcattctac ttaactttgc tcccagatgc ctccactct tcccagtact tgagaacaggaca ttcctcct tttaagcaca gactaaggct gtttatgtt ctctactcc ctcttcccac acaaccaaga gcctaaaggct gtttatgtt ctctactcc ctttcccac acaaccaccag gactaaggct cttcccct cataattcca gagaaaggca ttccttcct tttaagcaca gactaaggct gtttatgtt gtgcctcatt cctttcccac acaaccacaagaccac ctttccct cataattcca gagaaaggca ttccttcct accaccacaagcac gactaaggct gtttatgtt gtgcctcatt cctttcccac acaaccacaagaccacaagaccac accaccacaccacacaca	158 206 254 302 356 416 476 536 596 656 716 776 836 896 956
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly Als Strong 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac  Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20  25  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc  Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  35  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag  Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln  50  tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag  Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu  65  ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc geattctca  Leu Glu Val Asp Asp Trp Glu Phe  80  gccagggatg cagaggccac ccagaggccc ttcctgagg ccggccacat tcccgcctc  ctgggcagat tgggagata cttcactgct ccaaagcttt tgagacacaa gggaatctca  acaaccaggg atcaggagga tccaaagccg acattcccag tcctggagat tgggagatctca  acaaccaggg atcaggagga ccggcacct tcccaggagct tgactgctgg ctcattctaa  acaaccaggg atcaggagga ccgcacctct tcccaggagct tgactgcagc cccagggttt  ggctccttaa acccgaggac ccgcacctt tcccaggagct tgcaccagc ctcattctac  ttaactttgc tctcagatgc ctccagatgct ataggtcagt gaaagggca gtagtaagct  ggctcgctcc cttccgcag accttctccct cataattcca gagaaggca ttctctcct  cataattcc gagaaggca cttcctccct cataattcca gagaaggca ttctctctct	158 206 254 302 356 416 476 536 596 656 716 776 836 896

tcagagacgc aaaaaaaaaa aa

CCagagacge auducumum an	
<210> 104 <211> 346 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 170289	
<221> sig_peptide <222> 170250 <223> Von Heijne matrix	
<400> 104 ccatttgagc cccaccacgg aggttatgtg gtcccaaaag gaatgatggc caagcaatt	a 60
atttttctc ctagttctta gcttgcttct gcattgattg gctttacaca actggcatt agtctgcatt acacaaatag acactaattt atttggaaca agcagcaaa atg aga ac  Met Arg Th	t 178
tta ttt ggt gca gtc agg gct cca ttt agt tcc ctc act ctg ctt cta Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr Leu Leu -20 -15 -10	226
atc acc cct tct ccc agc cct ctt cta ttt gat aga ggt ctg tcc ctc  Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly Leu Ser Leu  -5 1 5	274
aga toa goa atg tot tagecootet cetetettee atteetteet gttggtacte Arg Ser Ala Met Ser	329
10 atttcttcta actttta	346
<210> 105 <211> 685 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 36497	
<pre>column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{column{colu</pre>	o
<221> polyA_site <222> 663685	
<pre>&lt;400&gt; 105 aagttctgcg ctggtcggcg gagtagcaag tggcc atg ggg agc ctc agc ggt</pre>	53
ctg cgc ctg gca gca gga agc tgt ttt agg tta tgt gaa aga gat gtt Leu Arg Leu Ala Ala Gly Ser Cys Phe Arg Leu Cys Glu Arg Asp Val	101
tcc tca tct cta agg ctt acc aga agc tct gat ttg aag aga ata aat Ser Ser Ser Leu Arg Leu Thr Arg Ser Ser Asp Leu Lys Arg Ile Asn	149

25 30 35	107
gga ttt tgc aca aaa cca cag gaa agt ccc gga gct cca tcc cgc act Gly Phe Cys Thr Lys Pro Gln Glu Ser Pro Gly Ala Pro Ser Arg Thr	197
tac aac aga gtg cct tta cac aaa cct acg gat tgg cag aaa aag atc Tyr Asn Arg Val Pro Leu His Lys Pro Thr Asp Trp Gln Lys Lys Ile	245
EE 60 65 70	293
ctc ata tgg tca ggt cgc ttc aaa aag gaa gat gaa atc cca gag act Leu Ile Trp Ser Gly Arg Phe Lys Lys Glu Asp Glu Ile Pro Glu Thr 75 80 85	
gtc tcg ttg gag atg ctt gat gct gca aag aac aag atg cga gtg aag Val Ser Leu Glu Met Leu Asp Ala Ala Lys Asn Lys Met Arg Val Lys 95	341
agc agc tat cta atg att gcc ctg acg gtg gta gga tgc atc ttc atg Ser Ser Tyr Leu Met Ile Ala Leu Thr Val Val Gly Cys Ile Phe Met 105 110 115	389
gtt att gag ggc aag aag gct gcc caa aga cac gag act tta aca agc Val Ile Glu Gly Lys Lys Ala Ala Gln Arg His Glu Thr Leu Thr Ser	437
ttg aac tta gaa aag aaa gct cgt ctg aaa gag gaa gca gct atg aag Leu Asn Leu Glu Lys Lys Ala Arg Leu Lys Glu Glu Ala Ala Met Lys	485
135 140 145 gcc aaa aca gag tagcagaggt atccgtgttg gctggatttt gaaaatccag	537
Ala Lys Thr Glu	597
gtatgatttg cccaaacctg taccatttcc gtatttctgc cgtagaagta gaaataaatt ttcttaaaaa aaaaaaaaa aaaaaaaa	657 685
ttottaaaaa aaaaaaaaaa uuuuuuu	
<210> 106 <211> 554	
<212> DNA	
<213> Homo sapiens	
<220> <221> CDS	
<222> 18320	
<221> polyA_signal	
<221> polyA_site <222> 542554	
<400> 106 aaccgtcgtg gggaagg atg gtg tgc gaa aaa tgt gaa aag aaa ctt ggt Met Val Cys Glu Lys Cys Glu Lys Lys Leu Gly 10	50
act grt arc act cca gat aca tgg aaa gat ggt gct agg aat acc aca	98
Thr Val Ile Thr Pro Asp Thr Trp Lys Asp Gly Ala Arg Ash Illi Illi 15 20 25	146
gaa agt ggt gga aga aag ctg aat aaa aat aaa gct ttg act tca aaa Glu Ser Gly Gly Arg Lys Leu Asn Lys Asn Lys Ala Leu Thr Ser Lys 30 35 40	
aaa gca aga ttt gat cca tat gga aag aat aag ttc tcc act tgt aga Lys Ala Arg Phe Asp Pro Tyr Gly Lys Asn Lys Phe Ser Thr Cys Arg	194
att tgt aaa agt tct gtg cac caa cca ggt tct cat tac tgc cag ggc Ile Cys Lys Ser Ser Val His Gln Pro Gly Ser His Tyr Cys Gln Gly	242
60 65 70 75  tgt gcc tac aaa aaa ggc atc tgt gcg atg tgt ggn aaa aaa gtt ttg	290

80 85 90	
gat acc aaa aac tac aag caa aca tct gtc tagatgtatt gatggaattt Asp Thr Lys Asn Tyr Lys Gln Thr Ser Val	340
ctggctttct aaatgatttt actttctgcc ttgaattttc aaggcataga tgtcaactta cagaataaca tgttttaaga taattaagtt taaaccagag aatttgattg ttactcattt tgctctcatg ttctaaacag caacagtgta actagtcttt tgttgtaaat ggttattttc cttataagaa ttttaagaac taaaaaaaaa aaaa	400 460 520 554
<210> 107 <211> 1678 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 711438	
<221> sig_peptide <222> 71136 <223> Von Heijne matrix score 3.5 seq AAPVAAGLGPVIS/RP	
<221> polyA_signal <222> 16441649	
<221> polyA_site <222> 16651678	
<400> 107 ccgacttcca gaggagcgct gtgcacgtgg agaagagcgg ggactcggcg accctgccct	60
ccgacttcca gaggagcgct gtgcacgtgg agaagagcgg ggactcggcg accetgcct cccgaccctc atg ttc gaa gag cct gag tgg gcc gag gcc cca gta Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val	60 1 <b>0</b> 9
ccgacttcca gaggagcgct gtgcacgtgg agaagagcgg ggactcggcg accetgcct cccgacctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20  -15  -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser	
ccgacttcca gaggagcgct gtgcacgtgg agaagagcgg ggactcggcg accetgcct cccgacctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20 -15 -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser  -5 1 5  tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg	109
ccgacttcca gaggagcgct gtgcacgtgg agaagagcgg ggactcggcg accetgcctt cccgaccttc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20  -15  -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser  -5  tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg  10  15  20  gcc cta gag gca gca tct ctt tcc cag cat ccc ccc agc cta tgt ata	109
ccgacttcca gaggagcgct gtgcacgtgg agaagagcgg ggactcggcg accetgcctt cccgaccctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20  -15  -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser  -5  tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg  10  15  gcc cta gag gca gca tct ctt tcc cag cat ccc ccc agc cta tgt ata Ala Leu Glu Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile  25  30  35	109 157 205
ccgacttcca gaggagcgct gtgcacgtgg agaagagcgg ggactcggcg accetgcctt cccgaccctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20  -15  -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser  -5  tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg  10  15  gcc cta gag gca gca tct ctt tcc cag cat ccc ccc agc cta tgt ata Ala Leu Glu Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile  25  agt gac tct gag gag gag gag gag gaa agg aag aag aa	109 157 205 253 301
ccgacctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20  -15  -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser  -5  tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg  10  gcc cta gag gca gca tct ctt tcc cag cat ccc ccc agc cta tgt ata Ala Leu Glu Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile  25  agt gac tct gag gag gag gag gag gaa agg aag aag aa	109 157 205 253
ccgacctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20  -15  -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser  -5  tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg  10  gcc cta gag gca gca tct ctt tcc cag cat ccc ccc agc cta tgt ata Ala Leu Glu Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile  25  agt gac tct gag gag gag gag gag gaa agg aag aag aa	109 157 205 253 301
ccgacctc atg ttc gaa gag cct gag tgg gcc gag ggactcggcg accetgcct cccgacctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta  Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val  -20  -15  -10  gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser  -5  tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg  10  gcc cta gag gca gca tct ctt tcc cag cat ccc ccc agc cta tgt ata Ala Leu Glu Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile  25  agt gac tct gag gag gag gag gag gaa agg aag aag aa	109 157 205 253 301

ata Ile	aat Asn	tca Ser	gcc Ala	cag Gln	cac His	ctg Leu	gac Asp	aat Asn	gtt Val	gac Asp	caa Gln	aca Thr	ggt Gly	ccc Pro	aaa Lys	541
120					125					130				•	135	
acc	taa	aag	ggt	agt	act	aca	aat	gat	cca	cca	aag	caa	agc	cct	ggg	589
Ala	Trp	Lys	Gly	Ser	Thr	Thr	Asn	Asp	Pro	Pro	Lys	Gln	Ser	Pro	Gly	
		•	-	140				-	145					150		
tcc	act	tee	cct		ccc	cct	cat	aca	tta	agc	cqc	aag	cag	tgg	cgg	637
502	Thr	Ser	Pro	Live	Pro	Pro	His	Thr	Leu	Ser	Arg	Lvs	Gln	Trp	Arg	
Ser	1111	261	155	Бys			****	160			5		165	•	_	
							252		224	326	220	+++		cca	cct	685
aac	cgg	caa	aag	aat	aag	aga	aya	Cor	aag	200	Tura	Dho	Cla	cca	Dro	
Asn	Arg		гàг	Asn	гув	Arg		Cys	гуу	ASII	nys		GIII	Pro	FIO	
		170					175					180				722
cag	gtg	cca	gac	cag	gcc	cca	gct	gag	gcc	ccc	aca	gag	aag	aca	gag	733
Gln	Val	Pro	Asp	Gln	Ala	Pro	Ala	Glu	Ala	Pro	Thr	Glu	Lys	Thr	GIu	
	185					190					195					
ata	tct	cct	qtt	ccc	agg	aca	gac	agc	cat	999	gct	cgg	gca	aāa	gct	781
Val	Ser	Pro	Val	Pro	Ara	Thr	Asp	Ser	His	Gly	Ala	Arg	Ala	Gly	Ala	
200		•			205		•			210		_			215	
++0	<b></b>	~~~	CGC	ata		cad	caa	cta	gat	aaa	acc	cga	ttt	cgc	tac	829
Log	724	712	720	Mot	Ala	Gln	Ara	Leu	Asp	GIV	Ala	Ara	Phe	Arg	Tyr	
ьęи	Arg	ATA	Arg		AIG	GIII	AL 9	200	225	017		9		230	- 2 -	
				220						-~+	aat	~~~	C 2 C		ctc	877
ctc	aat	gaa	cag	ttg	tac	tca	333	000	age	agt	37-	gca	Cag	cgt	Tou	0,,
Leu	Asn	Glu		Leu	Tyr	Ser	GIY		Ser	ser	Ala	Ala	GIN	Arg	neu	
		-	235					240					245			225
ttc	cag	gaa	gac	cct	gag	gct	ttt	ctt	ctc	tac	cac	cgc	ggc	ttc	cag	925
Phe	Gln	Glu	Asp	Pro	Glu	Ala	Phe	Leu	Leu	Tyr	His	Arg	Gly	Phe	GIn	
		250					255					260				
agc	caa	gtg	aag	aag	tgg	cca	ctg	cag	cca	gtg	gac	cgc	atc	gcc	agg	973
Ser	Gln	Val	Lys	Lys	Trp	Pro	Leu	Gln	Pro	Val	Asp	Arg	Ile	Ala	Arg	
	265		-	-	-	270					275					
gat		cac	cag	caa	cct	qca	tcc	cta	gtg	gtg	gct	gac	ttc	ggc	tgt	1021
Agn	Len	Ara	Gln	Ara	Pro	Ala	Ser	Leu	Val	Val	Ala	Asp	Phe	Gly	Cys	
280	1100	~- 9	022	••- 5	285					290		•		-	295	
200						+ < 2	aat	atc	caa		cct	ata	cat	tgc	ttt	1069
999	gat	Cyc	2	Tan	77-	Cox	Cor	Tle	722	Acn	Pro	Val	His	Cys	Phe	
GIY	ASD	Cys	AIG		Ala	Ser	Ser	110	305	1.011				310	•	
				300				~		~+~	tat	~ ~ ~	2 t C		cad	1117
gac	ttg	gct	tet	ctg	gac	200	agg	3703	mb-	919	Cura	) an	Mat	gcc	Gln	
Asp	Leu	Ala		Leu	Asp	Pro	Arg		THE	vai	Cys	Asp	Mec	Ala	G111	
			315					320					325			1165
gtt	cct	ttg	gag	gat	gag	tct	gtg	gat	gtg	gct	grg	דננ	tgc	ctt	CCa	1165
Val	Pro	Leu	Glu	Asp	Glu	Ser	Val	Asp	Val	Ala	Val		Cys	Leu	Sei	
		330					335					340				
ctg	atg	gga	acc	aac	atc	agg	gac	ttc	cta	gag	gag	gça	aat	aga	gta	1213
Leu	Met	Gly	Thr	Asn	Ile	Arg	Asp	Phe	Leu	Glu	Glu	Ala	Asn	Arg	Val	
	345					350					355					
cta	aaq	cca	aaa	aat	ctc	ctq	aaa	gtg	gct	gag	gtc	agc	agc	cgc	ttt	1261
Leu	Lvs	Pro	Glv	Glv	Leu	Leu	Lys	Val	Ala	Glu	Val	Ser	Ser	Arg	Phe	
360	-1-		1	2	365		•	•3		370					375	
	gat	att	cas	200		cta	caa	act	ata	acc	aad	cta	aac	ttc	aaq	1309
Glu	) and	Val	720	Thr	Dhe	Len	Ara	Ala	Val	Thr	Lvs	Leu	Glv	Phe	Lvs	
Giu	Asp	vai	Arg			пеа	2+3	n. u	385		-7.5	200	<b>4</b> -1	390	-2 -	
				380				200		++~	++~	++~			ttc	1357
att	gtc	ECC	aag	gac	ctg	acc	aac	age	TIL	Dho	Dho	Ton	Dhe	gat	Dhe	200.
ile	Val	Ser		Asp	Leu	Thr	Asn		HIS	Pne	Pne	neu	405	Asp	FIIC	
			395					400					405			1405
caa	aag	act	ggg	ccc	cct	ctg	gta	_ aāa	ccc	aag	gct	cag	CEE	tca	ggc	1405
Gln	Lys	Thr	Gly	Pro	Pro	Leu	Val	Gly	Pro	Lys	Ala			. Ser	Gly	
		410					415					420				
ctg	cag	ctt	cag	cca	tgt	ctc	tac	aag	cgc	agg	tga	cctc	tgg	atct	tccttg	1458
Leu	Gln	Leu	Gln	Pro	Cys	Leu	Tyr	Lys	Arg	Arg						
	425					430										
aga	gggg	agg	caga	tctc	aa a	ctcc	aggc	t ca	gaac	tgtg	aag	actg	ttt	ccgg	cctggc	1518
tgt	gago	caa	gacc	tggt	tc c	tggt	ggac	c ct	gagg	acaa	agt	gtga	taa	aacc	tctggc	1578
	_															

<210> 108	
<211> 494	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 25318	
<221> sig_peptide	
<222> 2575	
<pre>&lt;223&gt; Von Heijne matrix score 7.4</pre>	
seg FFLLLQFFLRIDG/VL	
•	
<221> polyA_signal	
<222> 452457	
<221> polyA_site	
<222> 482494	
<400> 108	51
aggctgagtg tgaagattag agta atg cct tct agc ttt ttc ctg ctg ttg  Met Pro Ser Ser Phe Phe Leu Leu	-
-15 -10	
cag tit the the aga att gat ggg gtg ett ate aga atg aat gae aeg	99
Gln Phe Phe Leu Arg Ile Asp Gly Val Leu Ile Arg Met Ash Asp Inc	
-5 1 5	147
aga ctt tac cat gag gct gac aag acc tac atg tta cga gaa tat acg	T. T.
Arg Leu Tyr His Glu Ala Asp Lys Thr Tyr Met Leu Arg Glu Tyr Thr	
to coa das ago ass att tot agt ttg atg cat gtt coa cot too cto	195
Ser Arg Glu Ser Lys Ile Ser Ser Leu Met His Val Pro Pro Ser Leu	
25 30 35 ⁴⁰	0.43
ttc acg gaa cct aat gaa ata tcc cag tat tta cca ata aag gaa gca	243
Phe Thr Glu Pro Asn Glu Ile Ser Gln Tyr Leu Pro Ile Lys Glu Ala	
45 50 55 gtt tgt gag aag cta ata ttt cca gaa aga att gat cct aac cca gca	291
Val Cys Glu Lys Leu Ile Phe Pro Glu Arg Ile Asp Pro Asn Pro Ala	
60 65 70	
gac tca caa aaa agt aca caa gtg gaa taaaatgtga tacaacatat	338
Asp Ser Gln Lys Ser Thr Gln Val Glu	
75 80	398
actcactatg gaatctgact ggacaccttg gctatttgta aggggttatt tttattatga gaattaatt	458
atogttacag goaggtttoa otoaaaaaaa aaaaac	494
staattsaa acsocttta ctcssssss sssssc	474

<210> 109 <211> 714 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 84..332

<221> sig_peptide <222> 84..170 <223> Von Heijne matrix score 5.2 seg PCYYLGLFQRALA/SV <221> polyA_site <222> 702..714 <400> 109 60 cctatctctt ctgctggctg ggctcaatgc cgcgggtgag cgttcggccg aggctgctcc taccettgag tgatgtgeet tga atg acg etg ett tea tte get get tte acg 113 Met Thr Leu Leu Ser Phe Ala Ala Phe Thr gct gct ttc tcc gtc ctc ccc tgt tac tac ctt ggg ctg ttt cag cgg 161 Ala Ala Phe Ser Val Leu Pro Cys Tyr Tyr Leu Gly Leu Phe Gln Arg -15 -10 geg etc geg teg gte tte gae eea ett tge gtt tgt tea egt gtg etc 209 Ala Leu Ala Ser Val Phe Asp Pro Leu Cys Val Cys Ser Arg Val Leu 257 ccg aca cct gta tgt acc ttg gtc gca aca caa gcc gaa aaa ata tta Pro Thr Pro Val Cys Thr Leu Val Ala Thr Gln Ala Glu Lys Ile Leu gag aat ggg ccc tgt cca acc aag gag gcg gcc cag ctt gtc ggg aag 305 Glu Asn Gly Pro Cys Pro Thr Lys Glu Ala Ala Gln Leu Val Gly Lys 40 352 ggc agc gtt tcc gcc aga aat gct tcg tgaaaggcac ttgagggacc Gly Ser Val Ser Ala Arg Asn Ala Ser 50 412 ttagcagcat cctcaacagg ccttgtaggg aatgccagaa gaagcagtcc ttggccgggc 472 ggggtggctc atgcctgtgg tcccagcact ttgggaggcc ggggcgggcg gatcacctga ggtcgggagg tccagaccag cctgaccgac atggagaaac cccgtctnta ctagaaatac 532 aaaactagcc gggtgtggtg gcgcatgcct gtagtcccag ctactcggga gggtgaggca 592 ggagacgttc ttgaacccgg gaggcggagt ttgtggtgag ccgagatcgc gccattgcac 652 712 714

<210> 110

<211> 805

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 32..718

<221> sig_peptide

<222> 32..100

<223> Von Heijne matrix score 7.4 seq VLLLAALPPVLLP/GA

<221> polyA_signal

<222> 770..775

<221> polyA_site

<222> 793..805

<400> 110

cctctttca	g cccgggat	cg ccccagca	gg g atg ggc Met Gly	gac aag atc tgg ctg Asp Lys Ile Trp Leu	52
ccc ttc c Pro Phe P -15	cc gtg ctc ro Val Leu	ctt ctg gc Leu Leu Al -10	c gct ctg co a Ala Leu Pr	-20 ot ccg gtg ctg ctg cct o Pro Val Leu Leu Pro -5	100
aga aca a	cc ggc tto la Gly Phe	aca cct tc	c ctc gat ag r Leu Asp Se 10	c gac ttc acc ttt acc r Asp Phe Thr Phe Thr 15	148
ctt ccc q	cc ggc cag	aag gag tg Lys Glu Cy	c ttc tac ca	g ccc atg ccc ctg aag n Pro Met Pro Leu Lys 30	196
Ala Ser L	tg gag ato	gag tac ca Glu Tyr Gl 40	n Val Leu As	t gga gca gga tta gat p Gly Ala Gly Leu Asp 45	244
att gat t	tc cat ctt	gcc tct cc	a gaa ggc aa	a acc tta gtt ttt gaa s Thr Leu Val Phe Glu 60	292
caa aga a	aa tca gat ys Ser Asp	gga gtt ca Gly Val Hi	c act gta ga s Thr Val Gl 75	g act gaa gtt ggt gat u Thr Glu Val Gly Asp 80	340
tac atg t	tc tgc ttt Phe Cys Phe	gac aat ac	a ttc agc ac r Phe Ser Th 90	c att tot gag aag gtg or Ile Ser Glu Lys Val 95	388
att ttc t Ile Phe P	tt gaa tta	atc ctg ga	t aat atg gg p Asn Met Gl 105	a gaa cag gca caa gaa y Glu Gln Ala Gln Glu 110	436
Gln Glu A	at tog aac	aaa tat at Lys Tyr Il 12	e Thr Gly Th	a gat ata ttg gat atg or Asp Ile Leu Asp Met 125	484
aaa ctg g	gaa gac ato	ctg gaa to Leu Glu Se 135	c atc agc ager Ile Ser Se	gc atc aag tcc aga cta er Ile Lys Ser Arg Leu 140	532
agc aaa a	agt ggg cad Ser Gly His	ata caa at	t ctg ctt ag e Leu Leu Ar. 15	ga gca ttt gaa gct cgt gg Ala Phe Glu Ala Arg 55 160	580
gat cga a	aac ata caa Asn Ile Glr 165	gaa agc aa Glu Ser As	ic ttt gat ag sn Phe Asp Ar 170	ga gtc aat ttc tgg tct rg Val Asn Phe Trp Ser 175	628
atg gtt a Met Val A	at tta gtg	gtc atg gt	g gtg gtg to al Val Val Se 185	a gcc att caa gtt tat er Ala Ile Gln Val Tyr 190	676
Met Leu L	ag agt cto	ttt gaa ga Phe Glu As 20	at aag agg aa p Lys Arg Ly	aa agt aga act 's Ser Arg Thr 205	718
taaaactcc	ca aactagag			gg cataaaaatg caataaactg	778 805

<210> 111

<211> 787

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 26..481

<221> sig_peptide

<222> 26..88

<223> Von Heijne matrix

score 4.4 seg AVASSFFCASLFS/AV

<221> polyA_signal <222> 755..760

<221> polyA_site <222> 775..787

<400> 111 gacagectgg ataaaggete acttg atg get eag ttg gga gea gtt gtg get Met Ala Gln Leu Gly Ala Val Val Ala gtg gct tcc agt ttc ttt tgt gca tct ctc ttc tca gct gtg cac aag 100 Val Ala Ser Ser Phe Phe Cys Ala Ser Leu Phe Ser Ala Val His Lys - 5 -10 ata gaa gag gga cat att ggg gta tat tac aga ggc ggt gcc ctg ctg Ile Glu Glu Gly His Ile Gly Val Tyr Tyr Arg Gly Gly Ala Leu Leu 15 10 act tog acc age ggc cot ggt tto cat oto atg oto cot tto atc aca 196 Thr Ser Thr Ser Gly Pro Gly Phe His Leu Met Leu Pro Phe Ile Thr 30 25 tca tat aag tot gtg cag acc aca ctc cag aca gat gag gtg aag aat 244 Ser Tyr Lys Ser Val Gln Thr Thr Leu Gln Thr Asp Glu Val Lys Asn 45 gta cct tgt ggg act agt ggt ggt gtg atg atc tac ttt gac aga att 292 Val Pro Cys Gly Thr Ser Gly Gly Val Met Ile Tyr Phe Asp Arg Ile 60 340 gaa gtg gtg aac ttc ctg gtc ccg aac gca gtg cat gat ata gtg aag Glu Val Val Asn Phe Leu Val Pro Asn Ala Val His Asp Ile Val Lys 75 aac tat act gct gac tat gac aag gcc ctc atc ttc aac aag atc cac Asn Tyr Thr Ala Asp Tyr Asp Lys Ala Leu Ile Phe Asn Lys Ile His 95 90 85 cac gaa ctg aac cag ttc tgc agt gtg cac acg ctt caa gag gtc tac 436 His Glu Leu Asn Gln Phe Cys Ser Val His Thr Leu Gln Glu Val Tyr 110 105 att gag ctg ttt gga ctg gaa aat gat ttt tcc cag gaa tct tca 481 Ile Glu Leu Phe Gly Leu Glu Asn Asp Phe Ser Gln Glu Ser Ser 125 120 taaaagggac cctgagcaag aacatttttc atagcagaca ggaggactca tccacatcgc 541 cagcaatcat aattaagcaa accgcctttt gcaccattta agatttagga aatcatccaa 601 attactttta atgtttctgc agtagaaaat gaatctaaat tcattttata gggtttgtag 721 tcttttatct gttttggatt cactgtgctt ttaagaaaaa gttggtaaat ttgccgttga tttttctttt taacctcaaa ctaatagaat tttataaaaat attaattttc tccaaaaaaa 781 787 aaaaaa

<210> 112

<211> 569

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 26..562

<221> sig_peptide

<222> 26..187

<223> Von Heijne matrix score 4.1

## seq AVVAAAARTGSEA/RV

		2														
<400	> 11	at c	tggg	ctac	a aa	agt	atg	gcc	gct	tct	gag	gcg	gcg	gtg	gtg	52
ayaa	acag	900	-555	-		- 3	Met	Ala	Āla	Ser	Glu	Ala	Ala	Val	Val	
											-50					7.00
tct	tcq	ccg	tct	ttg	aaa	aca	gac	aca	tcc	cct	gtc	ctt	gaa	act	gca	100
Ser	Ser	Pro	Ser	Leu	Lys	Thr	Asp	Thr	Ser	Pro	Val	Leu	Glu.	Thr	-30	
4 5					-40					-35						148
gga	acg	gtc	gca	gca	atg	gct	gcg	acc	ccg	tca	gca	agg	gct	gca N1a	712	140
Gly	Thr	Val	Ala	Ala	Met	Ala	Ala	Thr	Pro	ser	Ala	Arg	ALA	-15	AIG	
				-25					-20	<b>+</b>	~~~	acc	200		tcc	196
gcg	gtg	gtt	gcg	gcc	gcg	gcc	agg	acc	gga	505	Glu	Mla	Ara	Val	Ser	
Ala	Val	Val	Ala	Ala	Ala	Ala	Arg	-5	Gry	361	GIU	7.24	1			
			-10 ttg			224	cta		trc	tta	agc	aac	ata	ttc	gcc	244
aag	gcc	gct	Leu	gct Nla	Thr	Tive	Len	Leu	Ser	Leu	Ser	Gly	Val	Phe	Ala	
гÀг	AIA 5	Ala	теп	Ala	1111	10	200				15	•				
~+~	-	220	ccc	aaa	aaa	CCC	act	tca	gcc	gag	ctg	ctg	aat	cgg	ttg	292
val	Hie	Lvs	Pro	Lvs	Glv	Pro	Thr	Ser	Āla	Glu	Leu	Leu	Asn	Arg	200	
20					25					30					33	240
220	gag	aaq	ctg	ctg	gca	gaa	gct	gga	atg	cct	tct	cca	gaa	tgg	acc	340
Lys	Glu	Lys	Leu	Leu	Ala	Glu	Ala	Gly	Met	Pro	Ser	Pro	Glu	TIP	Thr	
_				40					45					50		388
aag	agg	aaa	aag	cag	act	ttg	aaa	att	999	cat	gga	999	act whr	Len	Asn	300
Lys	Arg	Lys	Lys	Gln	Thr	Leu	Lys	Ile	GIY	HIS	GIY	GIY	65	пец	rop	
			55					60	~~ ·	2++	aaa	agc		aca	aaa	436
agc	gca	gcc	cga Arg	gga	gtt	ctg	get	Val	Gly	Tle	Glv	Ser	Glv	Thr	aaa Lys	
Ser	Ala		Arg	GIY	vaı	Leu	75	Val	Gry	110	027	80	2		-	
	++~	70	201	ato	tta	tica		tcc	aaq	agg	tat	act	gcc	att	gga Gly	484
Mot	Ley	Thr	Ser	Met	Leu	Ser	Gly	Ser	Lys	Arg	Tyr	Thr	Ala	Ile	Gly	
	95					90					95					
gaa	a+ a	aaa	aaa	act	act	gat	aca	cta	gat	tct	acg	999	aag	gta	aca	532
Glu	Leu	Gly	Lys	Āla	Thr	Asp	Thr	Leu	Asp	Ser	inr	Gly	Lys	Val	1111	
100					105					110					115	569
gaa	gaa	aaa	cct	tac	ggt	ato	aac	cto	ato	: taa	gtag	Ī				505
Glu	Glu	Lys	Pro	Tyr	Gly	Met	Asr	Lev	ı Ile	<b>&gt;</b>						
				120					125							

<210> 113

<211> 893

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 4..810

<221> sig_peptide <222> 4..279

<223> Von Heijne matrix score 6.8 seq AVMLYTWRSCSRA/IP

X 4.

<221> polyA_signal

<222> 858..863

<221> polyA_site <222> 881..893

<40	0 > 1	13														
gcc						acc Thr										48
						ctc Leu										96
cca Pro	cct Pro -60	tcc Ser	tcc Ser	atc Ile	atg Met	tac Tyr -55	cag Gln	gct Ala	aac Asn	ttt Phe	gac Asp -50	aca Thr	aac Asn	ttt Phe	gag Glu	144
						acg Thr										192
aca Thr	gtc Val	cac His	tcc Ser	agc Ser -25	atg Met	aat Asn	gag Glu	atg Met	ctg Leu -20	gag Glu	gaa Glu	gga Gly	cat His	gag Glu -15	tat Tyr	240
						tgg Trp										288
						ccc Pro 10										336
gta Val 20	gag Glu	gtg Val	ctg Leu	gag Glu	ccg Pro 25	gag Glu	gtc Val	acc Thr	aag Lys	ctc Leu 30	atg Met	aag Lys	ttc Phe	atg Met	tat Tyr 35	384
ttt Phe	cag Gln	cgc Arg	aag Lys	gcc Ala 40	atc Ile	gag Glu	cgg Arg	ttc Phe	tgc Cys 45	agc Ser	gag Glu	gtg Val	aag Lys	cgg Arg 50	ctg Leu	432
						aag Lys										480
acc Thr	ctt Leu	ggc Gly 70	aag Lys	ttc Phe	atc Ile	aac Asn	atg Met 75	ttt Phe	gct Ala	gtc Val	ctg Leu	gat Asp 80	gag Glu	cta Leu	aag Lys	528
						aag Lys 90										576
gca Ala 100	cag Gln	ttc Phe	ctg Leu	cgg Arg	aag Lys 105	atg Met	gca Ala	gat Asp	ccc Pro	cag Gln 110	tct Ser	atc Ile	cag Gln	gag Glu	tcg Ser 115	624
cag Gln	aac Asn	ctt Leu	tcc Ser	atg Met 120	ttc Phe	ctg Leu	gcc Ala	aac Asn	cac His 125	aac Asn	agg Arg	atc Ile	acc Thr	cag Gln 130	tgt Cys	672
ctc Leu	cac His	cag Gln	caa Gln 135	ctt Leu	gaa Glu	gtg Val	atc Ile	cca Pro 140	ggc Gly	tat Tyr	gag Glu	gag Glu	ctg Leu 145	ctg Leu	gct Ala	720
gac <b>Asp</b>	att Ile	gtc Val 150	aac Asn	atc	tgt Cys	gtg Val	gat Asp 155	tac Tyr	tac Tyr	gag Glu	aac Asn	aag Lys 160	atg Met	tac Tyr	ctg Leu	768
act Thr	ccc Pro 165	agt Ser	gag Glu	aaa Lys	cat His	atg Met 170	ctc Leu	ctc Leu	aag Lys	gta Val	aaa Lys 175	ctc	ccc Pro	•		810
tga <u>c</u> ttta	gccg	Thr Pro Ser Glu Lys His Met Leu Leu Lys Val Lys Leu Pro 165 170 175  tgaggccgca cccatggagc ctgggcttac cctctcacct tcttcttatt aaaaatccgt tttaaaaaac aaaaaaaaa aaa												870 893		

<210> 114 <211> 1475 <212> DNA

<213> Homo sapiens

```
<220>
<221> CDS
<222> 55..459
<221> sig_peptide
<222> 55..120
<223> Von Heijne matrix
      score 7.2
      seq GLWLALVDGLVRS/SP
<221> polyA_signal
<222> 1444..1449
<221> polyA_site
<222> 1462..1475
<400> 114
cagtteegea getacgtgtg ggaccegetg etgateetgt egeagategt eete atg
                                                                     57
                                                           Met
105
Gln Thr Val Tyr Tyr Gly Ser Leu Gly Leu Trp Leu Ala Leu Val Asp
                        -15
ggg cta gtg cga agc agc ccc tcg ctg gac cag atg ttc gac gcc gag
                                                                     153
Gly Leu Val Arg Ser Ser Pro Ser Leu Asp Gln Met Phe Asp Ala Glu
                    1
atc ctg ggc ttt tcc acc cct cca ggc cgg ctc tcc atg atg tcc ttc
                                                                     201
Ile Leu Gly Phe Ser Thr Pro Pro Gly Arg Leu Ser Met Met Ser Phe
                                20
            15
atc ttc aac gcc ctc acc tgt gcc ctg ggc ttg ctg tac ttc atc cgg
                                                                     249
Ile Phe Asn Ala Leu Thr Cys Ala Leu Gly Leu Leu Tyr Phe Ile Arg
cga gga aag cag tgt ctg gat ttc act gtc act gtc cat ttc ttt cac
                                                                     297
Arg Gly Lys Gln Cys Leu Asp Phe Thr Val Thr Val His Phe Phe His
                        50
etc ctg ggc tgc tgg ttc tac agc tcc cgt ttc ccc tcg gcg ctg acc
                                                                     345
Leu Leu Gly Cys Trp Phe Tyr Ser Ser Arg Phe Pro Ser Ala Leu Thr
 tgg tgg ctg gtc caa gcc gtg tgc att gca ctc atg gct gtc atc ggg
                                                                     393 -
 Trp Trp Leu Val Gln Ala Val Cys Ile Ala Leu Met Ala Val Ile Gly
                                    85
                80
 gag tac ctg tgc atg cgg acg gag ctc aag gag ata ccc ctc aac tca
 Glu Tyr Leu Cys Met Arg Thr Glu Leu Lys Glu Ile Pro Leu Asn Ser
                                100
                                                                     489
 gcc cct aaa tcc aat gtc tagaatcagg ccctttggac atcccgctga
 Ala Pro Lys Ser Asn Val
 cacttgggcc ccttaacacc ttgggctgct cagaccctcc agatgaggtc cagcccagat
                                                                     549
 ctgagaggaa ccctggaaat gtgaagtctc tgttggtgtg ggagagatag tgagggcctg
                                                                     609
 tcaaagaagg caggtagcag tcagcatgac agctgcaaga atgacctctg tctgttgaag
                                                                     669
 cettggtate tgagaggtea ggaaggggae etetttgagg gtaataacat aattggaace
                                                                     729
 atgccactct tgagccacaa tacctgtcac cagcctgttg ttttaagaga gaaaaaaaat
                                                                     789
 caaggatate tgattggage aaaccaette tttagteate tgtettacet eeetgggaca
                                                                     849
 gctgttacct ttgcagtgtt gccgaatcac agcagttacc tttgcaatgt tgccgaatca
                                                                     909
                                                                     969
 cagcagttct gttggagaaa cgcttggttt ccggatccag agccacagaa agaaatgtag
 gtgtgaagta ttaggctgct gtcagggaga ggatggcaga tggaggcatc aagcacaagg
                                                                    1029
 aaaatgcaca acctgtgccc tgttatacac acgttcatgt gcgcccaaga acctatgact
                                                                    1149
 ttettecagt teettetace aggteceeat cetgetgeca geteteaaca tageaggeca
 taggacccag agaagaatcc cagtgttgct caaagtctga ccatcataaa gacactgcct
                                                                    1209
                                                                    1269
 gtcttctagg aatgaccagg cacccagctc ccactggact ccaatttttt ttcctgcctt
                                                                    1329
 atttagaatt ctttggcggg aagggtatga tgggttccca gagacaagaa gcccaacctt
                                                                     1389
 ctggcctggg ctgtgctgat agtgctgagg gagataggaa tttgctgcta agatttttct
```

ttggggtgga gtttcctctg tgaggggctt gcagctatcc ttcctgtgta tacaaataca

qtattttcca tgaaaaaaaa aaaaaa <210> 115 <211> 321 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 48..248 <221> sig_peptide <222> 48..161 <223> Von Heijne matrix score 6.3 seq LVFALVTAVCCLA/DG <221> polyA_signal <222> 283..288 <221> polyA_site <222> 308..321 <400> 115 gctgagaaga gttgagggaa agtgctgctg ctgggtctgc agacgcg atg aat aac 56 gtg cag ccg aaa ata aaa cat cgc ccc ttc tgc ttc agt gtg aaa ggc 104 Val Gln Pro Lys Ile Lys His Arg Pro Phe Cys Phe Ser Val Lys Gly -25 -30 cac gtg aag atg ctg cgg ctg gtg ttt gca ctt gtg aca gca gta tgc 152 His Val Lys Met Leu Arg Leu Val Phe Ala Leu Val Thr Ala Val Cys -10 -15 tgt ctt gcc gac ggg gcc ctt att tac cgg aag ctt ctg ttc aat ccc 200 Cys Leu Ala Asp Gly Ala Leu Ile Tyr Arg Lys Leu Leu Phe Asn Pro aac ggt cct tac cag aaa aag cct gtg cat gaa aaa aaa gaa gtt ttg 248 Asn Gly Pro Tyr Gln Lys Lys Pro Val His Glu Lys Lys Glu Val Leu 20 15 tgattttata ttacttttta gtttgatact aagtattaaa catatttctg tattcttcca 308 321 aaaaaaaaa aaa <210> 116 <211> 450 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 25..399 <221> sig_peptide <222> 25..186 <223> Von Heijne matrix

<400> 116

score 3.5

seq SILAQVLDQSARA/RL

ctgctccagc gctgacgccg agcc atg gcg gac gag gag ctt gag gcg ctg Met Ala Asp Glu Glu Leu Glu Ala Leu -50	51
agg aga cag agg ctg gcc gag ctg cag gcc aaa cac ggg gat cct ggt Arg Arg Gln Arg Leu Ala Glu Leu Gln Ala Lys His Gly Asp Pro Gly	99
gat gcg gcc caa cag gaa gca aag cac agg gaa gca gaa atg aga aac Asp Ala Ala Gln Gln Glu Ala Lys His Arg Glu Ala Glu Met Arg Asn -25 -20 -15	147
agt atc tta gcc caa gtt ctg gat cag tcg gcc cgg gcc agg tta agt Ser Ile Leu Ala Gln Val Leu Asp Gln Ser Ala Arg Ala Arg Leu Ser -10 -5 1	195
aac tta gca ctt gta aag cct gaa aaa act aaa gca gta gag aat tac Asn Leu Ala Leu Val Lys Pro Glu Lys Thr Lys Ala Val Glu Asn Tyr	243
ctt ata cag atg gca aga tat gga caa cta agt gag aag gta tca gaa Leu Ile Gln Met Ala Arg Tyr Gly Gln Leu Ser Glu Lys Val Ser Glu	291
caa ggt tta ata gaa atc ctt aaa aaa gta agc caa caa aca gaa aag Gln Gly Leu Ile Glu Ile Leu Lys Lys Val Ser Gln Gln Thr Glu Lys	339
aca aca aca gtg aaa ttc aac aga aga aaa gta atg gac tct gat gaa Thr Thr Thr Val Lys Phe Asn Arg Arg Lys Val Met Asp Ser Asp Glu	387
gat gac gat tat tgaactacaa gtgctcacag actagaactt aacggaacaa Asp Asp Asp Tyr	439
70 gtctaggaca g	450
<pre>&lt;210&gt; 117 &lt;211&gt; 1173 &lt;212&gt; DNA &lt;213&gt; Homo sapiens  &lt;220&gt; &lt;221&gt; CDS &lt;222&gt; 101137  &lt;221&gt; sig_peptide &lt;222&gt; 1072 &lt;223&gt; Von Heijne matrix</pre>	
<pre>&lt;400&gt; 117 gagctgctt atg gga cac cgc ttc ctg cgc ggc ctc tta acg ctg ctg Met Gly His Arg Phe Leu Arg Gly Leu Leu Thr Leu Leu Leu -20 -15 -10</pre>	51
-20 -15 -10  ccg ccg cca ccc ctg tat acc cgg cac cgc atg ctc ggt cca gag tcc  Pro Pro Pro Pro Leu Tyr Thr Arg His Arg Met Leu Gly Pro Glu Ser  -5 1 5	99
gtc ccg ccc cca aaa cga tcc cgc agc aaa ctc atg gca ccg ccc cga Val Pro Pro Pro Lys Arg Ser Arg Ser Lys Leu Met Ala Pro Pro Arg 10 15 20 25	147

	•															
atc Ile	gjy aaa	acg Thr	cac His	aat Asn	ggc Gly	acc Thr	ttc Phe	His	Cys	gac Asp	gag Glu	gca Ala	ctg Leu	gca Ala 40	tgc Cys	195
gca Ala	ctg Leu	ctt Leu	cgc Arg	30 ctc Leu	ctg Leu	ccg Pro	gag Glu	tac	35 cgg Arg	gat Asp	gca Ala	gag Glu	116	gtg	cgg Arg	243
	~~~	ant.	45	gaa Glu	a a a	ctc	act	50 tcc	tqt	gac	atc	gtg	gtg	gac	gtg	291
	~~~	60	+=C	G a C	cct	caa	65 aga	cac	cqa	tat	gac	cat	cac	cag	agg	339
Gly	Gly	Glu	Tyr	Asp	Pro	Arg 80	Arg	His	Arg	туг	Asp 85	urs	UIS	G111	72.5	387
Ser	Phe	Thr	Glu	acc Thr	Met 95	Ser	Ser	Leu	Ser	100	GTÅ	Arg	PIO	יבוב	105	
	aag Lys	ctg Leu	agc Ser	agt Ser	aca	gga Gly	ctc Leu	atc Ile	tat Tyr 115	ctg Leu	cac His	ttc Phe	ggg Gly	cac His 120	aag Lys	435
ctg Leu	ctg Leu	gcc Ala	Gln	110 ttg Leu	ctg Leu	ggc Gly	act Thr	agt Ser 130	gaa	gag Glu	gac Asp	agc Ser	atg Met 135	gtg	ggc Gly	483
acc Thr	ctc Leu	Tyr	125 gac Asp	aag Lys	atg Met	tat Tyr	gag Glu 145	aac	ttt Phe	gtg Val	gag Glu	gag Glu 150	gtg Val	gat Asp	gct Ala	531
gtg Val	Asp	Asn	gly ggg	atc Ile	tcc Ser	cag Gln 160	taa	gca Ala	gag Glu	Gly 999	gag Glu 165	. PLO	cga Arg	tat Tyr	gca Ala	579
Leu	Thr	act	acc Thr	ctg Leu	Ser	aca	cga Arg	gtt Val	gct Ala	cga Arg	Lev	aat	cct	acc Thr	tgg Trp 185	627
170 aac Asn		ccc	gac Asp	caa Gln	175 gac Asp	act Thr	gag Glu	gca Ala	999 Gly	ttc Phe	aac	cgt Arg	gca Ala	atg Met	, ADP	675
ctg Leu	gtt Val	caa Glr	ı Glu	Glu	ttt Phe	ctg Leu	cag Glr	n Arg	tta Leu	gat	tto Phe	tac Tyr	caa Glr 215	1111	agc Ser	723
tgg Trp	cts Lev	Pro	Ala		gcc	ttg Leu	ı val	. GIU	gac	g gco n Ala	ctt Lei	gcc Ala 230	caç Glr	g cga	ttc Phe	771
cag Glr	g gtg	220 g gad L Asp		a agt	gga Gly	Gli	1 I16	ato	gaa Glu	a cto 1 Leu	g gcg 1 Ala 24!	g aaa a Lys	ggt	gca / Ala	a tgt a Cys	819
Pro	Tr		g gag s Glu	g cat ı His	: Lev	туз	cad	c ctg s Lev	g gaa 1 Glu	a tci 1 Se: 26	gg Gl	g ctg	g tco ı Sei	c cct	c cca Pro 265	867
250 gtg Val		c at	c tto e Pho	e Phe	val	ato	tac Ty:	c act	gad Asj	c caq p Gl:	g gc	t gga a Gl	a cas	g tgg n Trj 28	g cga p Arg 0	915
ata Ile	a ca e Gl	g tg n Cy	s Va	l Pro	- 220	g gaq s Gl	g cc u Pr	c cad o His	tc s Se	a tt	c ca e Gl	a age	c cg r Ar 29	g ne	g ccc u Pro	963
ct; Le	g cc u Pr	o Gl	u Pr	a too	g cgg	g G1;	t ct y Le 30	t cgg	g ga	c ga p Gl	g gc u Al	c ct a Le	u As	c ca p Gl	g gtc n Val	1011
ag Se	r Gl	y Il	~ ~~	t gg o Gl	c tg y Cy	s Il	c tt e Ph	c at	c ca l Hi	t gc s Al	a ag a Se 32	I GI	c tt y Ph	c at e Il	t ggc e Gly	1059
Gl	y Hi		c ac	r Ar	g Gl	u Gl	t. ac	c tt a Le	g ag u Se	c at r Me	g go	c cg	t go g Al	c ac a Th	c ttg r Leu 345	1107
33 gc Al	c ca	g cg n Ar	jc to g Se	a ta r Ty	33 c ct r Le	c cc	a ca o Gl	a at n Il	c to e Se	c ta		aata	aaa	cctt		1157

350 355 1173

<210> 118
<211> 785
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 72..704
<221> sig_peptide
<222> 72..161
<223> Von Heijne matrix
score 13.2
seq_LLLLSTLVIPSAA/AP

<221> polyA_signal <222> 772..777

WO 99/31236

<400: cggaa agggg				~~~	722	acc	TCC	Pro	- 000	900		ggcg gca Ala	Arg	Arg	gaaga cat His	60 110
ctg ( Leu )	Leu	Val	Leu	Leu	Leu	Leu	Leu -10	ser	Thr	Dea	Vai	-5	110	tcc Ser		158
gca Ala	gct Ala	cct Pro	Ile	His	Asp	Ala	gac Asp	Ala	GIII	10	361				15	206
aca Thr	Gly	Leu	Gln	Ser	cta Leu	Leu	GIN	GIY	25	261	Arg	ctt Leu	2 1.0	30	-1 -	254
ggt Gly	aac Asn	ctg Leu			ggc Gly	ata Ile	gac Asp	agc Ser 40	tta Leu	ttc Phe	tct Ser	gcc Ala	CCC Pro 45	atg Met	gac Asp	302
Phe	Arg	Gly	ctc Leu	Pro	Gly	Asn	Tyr	HIS	гуя	GIU	GIU	aac Asn 60	0111	010		350
cag Gln	Leu	gly ggg 50	aac Asn	aac Asn	acc Thr	ctc Leu 70	tcc	agc Ser	cac His	ctc Leu	cag Gln 75	atc Ile	gac Asp	aag Lys	gta Val	398
Pro	65 agg Arg	atg Met	gag Glu	gag Glu	aag Lys 85	a=a	gcc Ala	ctg Leu	gta Val	ccc Pro 90	atc Ile	cag Gln	aag Lys	gcc Ala	acg Thr 95	446
80 gac Asp	agc Ser	ttc Phe	cac His	Thr	<b>~</b> 22	ctc Leu	cat His	ccc Pro	cgg Arg 105	qtq	gcc Ala	ttc. Phe	tgg Trp	atc Ile 110		494
aag Lys	ctg Leu	cca Pro	Arg	100 cgg Arg	agg Arg	tcc Ser	cac His	cag Gln 120	gat Asp	gcc Ala	ctg Leu	gag Glu	ggc Gly 125	1	cac His	542
tgg Trp	ctc <b>Le</b> u	Ser	Glu	aag Lys	cga Arg	cac His	cgc Arg 135	ctg Leu	cag	gcc Ala	atc Ile	cgg Arg 140	rop	gga Gly	ctc	590
cgc Arg	Lys	Gly		cac His	aag Lys	Asp	gtc Val	cta	gaa Glu	gag Gļu	999 Gly	acc Thr	gag	ago Ser	tcc Ser	638
tcc	145 cac	tcc	agg:	ctg	tcc	150	cga	aag	acc	cac			tac	ato	ctc	686

	His :	Ser I	Arg I	Leu	Ser 165	Pro I	Arg 1	Lys :	Thr	His I 170	Leu 1	Leu'	Tyr :	lle :	Leu 175 -	
160 agg Arg	ccc Pro	tct Ser	Arg (	cag Gln	ctg	tagg	ggtg	gg g	accg	gggag	g ca	cctg	cctg	٠		734
tago	cccc	at c	agaco	180 cctg	c cc	caag	cacc	ata	tgga	aat a	aaag	ttct	tt c			785
<211 <212	)> 11 .> 55 ?> DN 3> Ho	9 A	apie	ns												
	)> L> CD 2> 44		5													
<223 <223	se	on He core eq LV	3 ijne	mat		/wm										
<40	0> 13 aacca	19 aga g	gggag	atga	at ca	acctg	gaaco	act	gcto	ccaa	acc	atg Met -60	ggc	agt Ser	aaa Lys	55
tgc Cys	tgt Cys -55	aaa Lys	ggt Gly	ggt Gly	cca Pro	gat Asp -50	gaa Glu	gat Asp	gca Ala	gta. Val	gaa Glu -45	aga Arg	cag Gln	agg Arg	cgg Arg	103
Gln	aag Lys	ttg Leu	ctt Leu	ctt Leu	gca Ala -35	caa	ctg Leu	cat His	cac His	aga Arg -30	aaa Lys	agg Arg	gtg Val	aag Lys	gca Ala -25	151
-40 gct Ala	Gly 999	cag Gln	atc Ile	cag Gln -20	acc	tgg Trp	tgg Trp	cgt Arg	999 Gly -15	gtc Val	ctg Leu	gtg Val	cgc <b>Arg</b>	agg Arg -10	acc Thr	199
ctg Leu	ctg Leu	gtt Val	gct Ala -5	acc	ctc Leu	agg Arg	gcc Ala	tgg Trp 1	atg Met	att Ile	cag Gln	tgc Cys 5	tgg Trp	tgg Trp	agg Arg	247
acg Thr	ttg Leu 10	gtg ·Val	cad	Arg	cgg Arg	Ile	cgt Arg	cag Gln	cgg Arg	cgg Arg	cag Gln 20	gcc Ala	ctg Leu	ttg Leu	agg Arg	295
gto Val 25	tac Tyr	gtc Val	atc Ile	cag	gag	caq	gcg Ala	acg Thr	gtc Val	aag Lys 35	ctc Leu	cag Gln	tcc Ser	tgc Cys	atc Ile 40	343
cac	atg Met	tgg Trp	cag Gln	tgc Cys 45	caa	caa Gln	tgt Cys	tac Tyr	cgc Arg 50	caa Gln	atg Met	tgc Cys	aat Asn	gct Ala 55	ctc Leu	391
tgo Cys	ttg Leu	ttc Phe	cag Gln 60	ato	cca	gag Glu	agc Ser	agc Ser 65	ctt	gcc Ala	ttc Phe	cag Gln	act Thr 70	gat Asp	ggc	439
ttt Phe	tta e Leu	Gln	atc	caa Glr	tat Tyr	gca Ala	atc Ile 80	cct	tca Ser	aag Lys	cag Gln	cca Pro	gag Glu	ttc Phe	cac His	487
	t gaa							rcct	<b>3</b> 999	gcatg	ga g		ggct	g		535
ca	90 ctacc	cta	ataa	atgt	ct g	gacc										559

```
<210> 120
<211> 770
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 25..393
<221> sig_peptide
<222> 25..150
<223> Von Heijne matrix
      score 4.6
      seq LDPAVSLSAPAFA/SA
<221> polyA_signal
<222> 734..739
<221> polyA_site
<222> 757..770
<400> 120
cgcagaaagg agagacacac atac atg aaa gga ggt ttc tcc aat ctt
                                                                       51
                           Met Lys Gly Gly Ala Phe Ser Asn Leu
                                   -40
aat gat tcc cag ctc tca gcc tcg ttt ctg caa ccc agc ctg caa gca
Asn Asp Ser Gln Leu Ser Ala Ser Phe Leu Gln Pro Ser Leu Gln Ala
                                -25
                                                    -20
            -30
aac tgt cct gct ttg gac cct gct gtg tca ctc tcc gca cca gcc ttt
                                                                      147
Asn Cys Pro Ala Leu Asp Pro Ala Val Ser Leu Ser Ala Pro Ala Phe
                                                -5
                            -10
gcc tct gct ctt cgc tct atg aag tcc tcc cag gct gca cgg aag gac
                                                                      195
Ala Ser Ala Leu Arg Ser Met Lys Ser Ser Gln Ala Ala Arg Lys Asp
                   - 5
gac ttt ctc agg tct ctt agt gat gga gac tca ggg aca tca gaa cac
                                                                      243
Asp Phe Leu Arg Ser Leu Ser Asp Gly Asp Ser Gly Thr Ser Glu His
                                    25
                20
atc tca gcg gtg gtg act agc cct cgg att tcc tgc cat ggt gct gcc
                                                                      291
Ile Ser Ala Val Val Thr Ser Pro Arg Ile Ser Cys His Gly Ala Ala
                                40
                                                    45
                                                                      339
att ccc acc gcc cgt gcc ctc tgc cta ggc tgt tcc tgc tgc acc gaa
Ile Pro Thr Ala Arg Ala Leu Cys Leu Gly Cys Ser Cys Cys Thr Glu
                            55
                                                60
                                                                      387
cgc ctc ctc ctg cca ccg ccc tcc ctc ctt tct tta gaa gcc cct gcc
Arg Leu Leu Pro Pro Pro Ser Leu Leu Ser Leu Glu Ala Pro Ala
                        70
                                                                      443
age ace tgagetetet getgattget gtteeteeca gtetgtggaa getttgeeca,
Ser Thr
tatgctttcc ttaaaagggt tctgggcagg gcaggcgccc ccatttctca gggatcccct
                                                                      503
ccaggacaac gccttttcct tgtgtcttca gctctcctta ccagatatct atatatttgt
                                                                      563
                                                                      623
atatattcag tttcaccaac aatgcatcaa gtactttttt ttttaagtaa agaaccgcag
tcatcgaact ggagccccat tgattccctc cccctcgcct ccccaaatct ggcacctgcc
                                                                      683
                                                                      743
caaggtatcc tcagaaccat ttggggtgtc ctttggcatt ggataataga aataaaattt
                                                                      770
tacctctttc tacaaaaaaa aaaaaac
```

<210> 121

<211> 1213

<212> DNA

<213> Homo sapiens

<220>																•
<221>	CDS	}												•		
<222>			95													
<221>				•												
<222>	· Jo. · Vor	He	: iine	mati	cix											
<b>4223</b>	SCC	re !	5.4													
	sec	LSI	HLLPS	SLRQ\	/IQ/E	EP										
<221: <222:																
<400	> 12	1									·	~~	aa a	2020	CC.	57
cctg	gctt	tg c	cttt	gccc	t gc	tgtg	tgat	ctt	agct	ccc	tgcc	cagg	cta	caa	caq	105
atg	gcc	atg .	gcc	cag	aaa	CCC i	agc Ser	His	Leu	Leu	Pro	Ser :	Leu .	Arg	Gln	
				. 7 🖺					-10					-		
gtc	atc	caq			cag	cta	tct	ctg	cag	cca	gag	cct	gtc	ttc	acg	153
Val	Ile	Gln	Glu	Pro	Gln	Leu	Ser	Leu	Gln	Pro	GIU	FIU	Val	Pne	Thr	
			-				<b>—</b>					<b>+</b> •				201
gtg	gat	cga	gct	gag Glu	gtg	cca	ecg Pro	Len	Phe	Tro	Lvs	Pro	Tyr	Ile	Tyr	
						20					42					
	15	tac	caa	ccg	a+a	cat	cag	acc	tgg	cgc	ttc	tat	ttc	cgc	acg	249
Ala	Glv	Tvr	Arg	Pro	Leu	His	Gln	Thr	Trp	Arg	Phe	Tyr	Phe	Arg		
					3 =					40						297
ctg	ttc	cag	cag	cac	aac	gag	gcc	gtg	aat	gtc	tgg	Thr	His	Leu	Leu	
Leu	Phe	Gln	Gln	His	Asn	GIU	Ala	vai	55	vaı	111			60		
			~+ >	50 ctg	cta	cta	caa	cta	acc	ctc	ttt	gtg	gag	acc	gtg	345
gcg	gcc	Leu	Val	Leu	Leu	Leu	Arg	Leu	Ala	Leu	Phe	Val	Glu	Thr	Val	
			C =					70								202
gac	ttc	tgg		gac	cca	cac	gcc	ctg	CCC	ctc	ttc	atc	att	gtc	CEE	393
Asp	Phe	Trp	Gly	Asp	Pro	His	Ala	Leu	Pro	Leu	Phe	30 11e	TTE	Val	пец	
		~ ~					85					<i>-</i>				441
gcc	tct	ttc	acc	tac Tyr	CEC	Ser	Len	Ser	Ala	Leu	Ala	His	Leu	Leu	Gln	
	~ -					ากก					TOD					
acc		tct	gag	ttc	tgg	cat	tac	agc	ttc	ttc	ttc	ctg	gac	tat	gtg	489
Ala	Lys	Ser	Glu	Phe	Trp	His	Tyr	Ser	Phe	PHE	Phe	Leu	Asp	Tyr	Val 125	
					775					120						537
999	gtg	gcc	gtg	tac	cag	ttt	ggc	agt	gcc	Ley	gca	His	Phe	Tvr	Tyr	
Gly	Val	Ala	Val	Tyr	GIn	Pne	GIY	Ser	135	рец	лта			140	•	
	250	~ ~ ~		130	+ ~~	cat	acc	caq	ata	cæa	gct	gtt	ttt	ctg	ccc	585
gct Ala	Tle	Glu	Pro	Ala	Tro	His	Ala	Gln	Val	Gln	Ala	Val	Phe	Leu	Pro	
			115					150					100			633
atg	gct	gcc	ttt	ctc	gcc	tgg	ctt	tcc	tgc	att	ggc	tcc	tgc	tat	aac	633
Met	Ala	Ala	Phe	Leu	Ala	Trp	Leu	Ser	Cys	; Ile	GIY	170	د ړ د	TAT	Asn	
		160	)				165			· car	· aca			gac	gtg	681
aag	tac	ato	cag	aaa	CCa	ggc	Len	Leu	Glv	/ Arc	Thr	Cys	Gln	Glu	g gtg val	
	376					ารถ	1				703	,				
ccc			cto	acc	tac	aca	cto	gac	att	agt	cct	gtg	gtg	cat	cgt Ara	729
Pro	Ser	· Val	Lev	Ala	туг	Āla	Let	ı Asp	ıle	e ser	PIC	val	. Val	His		
101	١				195	;				200	,					777
ato	tto	gt	tco	tcc	gac	; ccc	acc	acc	gat	gat	, CCS	a gct	Lei	. Lei	tac ı Tyr	
				210	•				21!	5				22		
C = 1	- 213/	ta.	c cas	zito ato	, a ato	: tto	tt!	cte	g ct	g gci	t gct	gc	tto	: tt	c tct	825
Cal	- 44	اوع و		, ,,,,,	, ,,,,,,,			•	- '		-					

His Lys Cys Gln Val Val Phe Phe Leu Leu Ala Ala Ala Phe Phe Ser	
225 230 230 ago too too too ggg ago too cat gto tto ggg	873
Thr Phe Met Pro Glu Arg Trp Phe Pro Gly Ser Cyo Mile 1250	
and the the cat are the tra gtg etg tge acg etg get	921
Gln Gly His Gln Leu Phe His He Phe Leu var heu cys 1112 265	060
and get get get gag tat gag gec ega egg ecc atc tat	969
Gln Leu Glu Ala Val Ala Leu Asp Tyr Glu Ala Alg Alg 110 210 210 285	
270 270 and and true cot can each tit tot age of the ctg	1017
Glu Pro Leu His Thr His Trp Pro His Ash Phe Ser Gly 200 100	
290 295  ctc acg gtg ggc agc agc atc ctc act gca ttc ctc ctg agc cag ctg  ctc acg gtg ggc agc agc atc ctc act gca ttc ctc ctg sec cag ctg	1065
Leu Thr Val Gly Ser Ser Ile Leu Thr Ala Phe Leu Leu Ser Gln Leu	
305 310	1115
gra cag cgc aaa ctt gat cag aag acc aag tgaaggggga tggcatctgg	1110
Val Gln Arg Lys Leu Asp Gln Lys Thr Lys 320 325	
taggagga ggtatagttg ggggacaggg gtctgggttt ggctccaagt gggaacaagg	1175
cctggtaaag ttgtttgtgt ctggccaaaa aaaaaaaa	1213
<210> 122	
<211> 1318	
<212> DNA <213> Homo sapiens	
CZ137 Hollo Daptolla	
<220>	
<221> CDS <222> 31660	
<2222> 31000	
<221> sig_peptide	
<222> 3190	
<223> Von Heijne matrix score 5.4	
seq AFVIACVLSLIST/IY	
one welch signal	
<221> polyA_signal <222> 12881293	
<221> polyA_site	
<222> 13071318	
<400> 122	54
ggaggatggg cgagcagtct gaatgccaga atg gat aac cgt ttt gct aca gca Met Asp Asn Arg Phe Ala Thr Ala	
-20 -15	
ttt gta att gct tgt gtg ctt agc ctc att tcc acc atc tac atg gca	102
Phe Val Ile Ala Cys Val Leu Ser Leu IIe Ser In Ile 171 100 1100	
-10	150
Ala car the cly Thr Ash Phe Tro TVI Glu IVI AIG Ser 110 tall of	
E 10 15 T	198
gaa aat too agt gat tog aat aaa ago ato tgg gat gaa tto att agt	
Glu Asn Ser Ser Asp Leu Asn Lys Ser Ile Trp Asp Glu Phe Ile Ser	
and and and act tat aat gat gca ctt ttt cga tac aat	246
Asp Glu Ala Asp Glu Lys Thr Tyr Ash Asp Ala Bed File 150	
40 45	

ggc Gly	aca Thr	Val	gga Gly	ttg Leu	tgg Trp	aga Arg	Arg	tgt Cys	atc Ile	acc Thr	ata Ile	ccc Pro 65	aaa Lys	aac Asn	atg Met	294
cat His	tgg Trp	55 tat Tyr	agc Ser	cca Pro	cca Pro	Glu	60 agg Arg	aca Thr	gag Glu	tca Ser	FIIC	gat	gtg Val	gtc Val	aca Thr	342
	70	~+~	201	++0	202	75 cta	act	gag	caq	ttc	atg	gag	aaa	ttt Phe	gtt Val	390
85		<b>~</b> ~~	330	Cac	90 aat	agc	aaa	att	gat	ctc	ctt	agg	acc	tat	ctt	438
				105	ctt	tta	cct	ttt	ata	aqt	tta	ggt	ttg	Tyr 115 atg	tgc	486
Trp	Arg	Cys	Gln	Phe	Leu	Leu	Pro	125	val	Ser	теп	GIY	130	Met tta	0,70	534
Phe	Gly	Ala	,Leu	Ile	Gly	Leu	Cys	АТа	Cys	116	Cys	145	361	200	-1-	500
ccc Pro	Thr	att Ile	gcc Ala	acg Thr	ggc Gly	Ile	ctc Leu	cat His	ctc Leu	ctt Leu	gca Ala 160	AGT	aca Thr	aag Lys	gag Glu	582
agc Ser	150 atg Met		cca Pro	gct Ala	gga Gly	155 gct Ala	gag Glu	tcc	aag Lys	HIS	aca Thr	gcc	act Thr	cct Pro	gca Ala 180	630
165 cac	gca	tac	gta		170 aca	ggg	aag	ccc	aag	tag				agag		680
				105					190	)	aaa	gaca	aat	ctac	ttccct	740
_				. ~ + ~ +	30 t	+	CCCT	c ac	Jaact	Caaa	lace	aatt	.904			800
+			+ ~++		ac t	gaat	atqa	וב כז	:cca:	lacco	:				, = = = 5 -	860
			~+ ~ =	taas	CO 2	rtoca	ıacca	la Ca	igacc	gcaat		Lyas	gaay	acgo		920 980
		-+-	++at		itt c	racct	aact	a a	actca	ggaa	Lace	2995	gaag	augi		1040
- ~ -	+	+	aato	1+222	at t	せたたた	ctac	it ta	aaato	ctatt	: tat	こしししし		gray	19005	1100
	-++-1	+ ~ ~	cact		ot c	retet	aato	at at	caaca	aaacc	: 99:	.caac	lacc			1160
cct	cct	gata	gtag	gttga	at o	ctac	CCCC	gc av	tacci	.aacc	y cat	1000	aaaa	aatt	atctag ctgtat	1220
cag	gaaat	caca	caco	ccca	aaa a	acaca	·++a	-t ti	+++-	taac	, 500	actat	ata	ttga	ctgtat	1280
acq	gaati	tatt	aaai	ttati	cc t	tctg	ggaaa	aa a	aaaa	aaa				-		1318

<211> 853

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 31..582

<221> sig_peptide

<222> 31..90

<223> Von Heijne matrix score 5.4 seq AFVIACVLSLIST/IY

<221> polyA_signal

<222> 816..821

<221> polyA_site <222> 840..853

								- 20		•	_		gct Ala -15			54 102
Phe	Val	Ile	Ala	Cys	Val	цеu	267	ctc Leu	att Ile			1	tac Tyr			150
Ala	Ser	Ile	Gly	Thr	Asp	Pne	тър	- y -		15	_		cca Pro		20	
5 gaa Glu	aat Asn	tcc Ser	agt Ser	gat Asp	10 ttg Leu	aat Asn	aaa Lys	agc Ser		tgg Trp	gat Asp	gaa Glu	ttc Phe	att Ile 35	agt Ser	198
				25					73±	aca	cct	ttt	cga Arg 50	tac	aat	246
			40					4	-+-	200	ata	ccc	aaa Lys	aac	atg	294
		55							as a	tca	ttt	gat	gtg Val	gtc	aca	342
	70					/5			~~~	ttc	ato	gag	aaa Lys	ttt	gtt	390
85					90				gat Asp	cto Lev	ctt	agg	acc	tat	ctt Leu	438
				105	•			ttt Phe	gtg Val	, ant	- ++=	a aat	ttq	atg	tgc Cys	486
			120	)				- ~~	, - +a	~ att	ta	c cqa	a ago	tta	a tat ı Tyr	534
		13	5				T4.		- c+	c ct	t gc	a gat a Asj	t acc	ate	g ctg t Leu	582
F 1.	7.5	^	•			15	5				16	0			~==	642
		~ ~~~~	444	acat	aga i	aata	tcct	gt g	taga	tgct	c ca	gctg	aaat	CCC	aagctaa atgtcca	702
tg	aagt	ccay	900	2000	99C	atca	tttc	ca g	ccat	gtgt	g 99	agcc	atcc	tgg	atgtcca ttgtgag	762
go	CCCC	aact	gac	4460	aaa	gact	tcaq	cc a	cagc	tatt	a tc	ttac	taca	tcc	ttgtgag aataaaa	822
20	ctta tcta aatt	ataa	aga	acca	act	agcı	gage	CC a	a	acct	a tg	gaac	tgat	aga	aataaaa	822 853

<211> 826

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 15..695

<221> sig_peptide <222> 15..80 <223> Von Heijne matrix score 8.5 seq AALLLGLMMVVTG/DE

<221> polyA_signal <222> 795..800

<221> polyA_site <222> 814..826

<400> 124				
aaccagaggt gccc	atg ggt tgg	aca atg agg	ctg gtc aca gca	gca ctg 50
		Thr Met Arg	Leu Val Thr Ala	Ala Leu
	-20		-15	
tta ctg ggt ctc	atg atg gtg	gtc act gga	gac gag gat gag	aac agc 98
		Val Thr Gly	Asp Glu Asp Glu	Asn Ser
-10	-5		I	tac caa 146
ccg tgt gcc cat	gag gcc ctc	ten Arm Glu	gac acc ctc ttt	-5
pro cys Ala HIS	GIU AIA LEU	15	Asp Thr Leu Phe 20	cys cin
	ttc tac cca		aac att ggc tgc	aag gtt 194
Gly Len Glu Val	Phe Tvr Pro	Glu Leu Glv	Asn Ile Gly Cys	Lys Val
25		30	35	•
	aac aac tac	aga cag aag	atc acc tcc tgg	atg gag 242
Val Pro Asp Cys	Asn Asn Tyr	Arg Gln Lys	Ile Thr Ser Trp	Met Glu
40	45		50	
ccg ata gtc aag	ttc ccg ggg	gcc gtg gac	ggc gca acc tat	atc ctg 290
Pro Ile Val Lys		Ala Val Asp	Gly Ala Thr Tyr	
55	60		65	70
gtg atg gtg gat	cca gat gcc	cct agc aga	gca gaa ccc aga	cag aga 338
Val Met Val Asp		Pro Ser Arg	Ala Glu Pro Arg	85
tta taa 252 52t	75		aag ggc gcc gac	
Dhe Tro Ara His	Trn Leu Val	Thr Asp Ile	Lys Gly Ala Asp	5
90	ith nea var	95	100	
	cag ggc cag		gcc tac cag gct	ccc tcc 434
Lvs Glv Lvs Ile	Gln Gly Gln	Glu Leu Ser	Ala Tyr Gln Ala	Pro Ser
105		110	115	
cca ccg gca cac	agt ggc ttc	cat cgc tac	cag ttc ttt gtc	tat ctt 482
Pro Pro Ala His	Ser Gly Phe	His Arg Tyr	Gln Phe Phe Val	Tyr Leu
120	125		130	
cag gaa gga aag	gtc atc tct	ctc ctt ccc	aag gaa aac aaa	act cga 530
		Leu Leu Pro	Lys Glu Asn Lys	150
135	140	+++ c+c >>c	145	
ggc tet tgg aaa	Mot Non Nra	Dhe Leu Asn	cgt ttc cac ctg Arg Phe His Leu	33-3
Già sei iib mas	155	160	Arg riic iiib bea	165
cct gaa gca agc			aac tac cag gac	
Pro Glu Ala Ser	Thr Gln Phe	Met Thr Gln	Asn Tyr Gln Asp	Ser Pro
170		175	180	
acc ctc cag gct	ccc aga gaa	agg gcc agc	gag ccc aag cac	aaa aac 674
Thr Leu Gln Ala	Pro Arg Glu	Arg Ala Ser	Glu Pro Lys His	Lys Asn
185		190	195	
			getttgecat ceggg	catgt 725
Gln Ala Glu Ile	_			
200	205			cagaacccct 785
			accc cctctggata	cagaaccccc 765 826
tcttttccaa ataaa	aaaaaa aatca	cccaa aaaaaa	aaad d	520

<210> 125

<211> 571

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 74295	
<221> sig_peptide <222> 74196 <223> Von Heijne matrix	
<221> polyA_signal   <222> 545550	
<221> polyA_site <222> 561571	
<pre>&lt;400&gt; 125 cgggtagtgg tcgtcgtggt tttccttgta gttcgtggtc tgagaccagg cctcaagtgg aaacggcgtc acc atg atc gca cgg cgg aac cca gta ccc tta cgg ttt</pre>	60 109
ctg ccg gat gag gcc cgg agc ctg ccc ccg ccc aag ctg acc gac ccg Leu Pro Asp Glu Ala Arg Ser Leu Pro Pro Pro Lys Leu Thr Asp Pro	157
cgg ctc ctc tac atc ggc ttc ttg ggc tac tgc tcc ggc ctg att gat Arg Leu Leu Tyr Ile Gly Phe Leu Gly Tyr Cys Ser Gly Leu Ile Asp	205
aac ctg atc cgg cgg agg ccg atc gcg acg gct ggt ttg cat cgc cag Asn Leu Ile Arg Arg Arg Pro Ile Ala Thr Ala Gly Leu His Arg Gln 5 10 15	253
ctt cta tat att acg gcc ttt ttt ttg ctg gat att atc ttg Leu Leu Tyr Ile Thr Ala Phe Phe Leu Leu Asp Ile Ile Leu 20 25 30	295
taaaacgtga agactacctg tatgctgtga gggaccgtga aatgtttgga tatatgaaat tacatccaga ggattttcct gaagaagata agaaaacata tggtgaaatt tttgaaaaat tccatccaat acgttgaagt cttcaaaatg cttgctccag tttcactgat acctgctgtt cctgaatttg atggaacatg tttcttatga cagttgaagc ttatgctaat ctgtatgttg acaccttgta attaaaatac gtaccaaaaa aaaaaa	355 415 475 535 571
<210> 126 <211> 659 <212> DNA <213> Homo sapiens	
<220> <221> CDS o<222> 440658	
<221> polyA_signal <222> 601606	
<pre>&lt;400&gt; 126 cgccttacga gctgggaggt ggtgcctctc acccagctaa ttgctctcta gcccttggcc ttcacaggtg ttggtgcctg ccgtgaacgc attctgacct gggccgtatc tgtctccaa gactttgtgc ctatggttgg ggacagagtg aggtcgttgc cttgacgacg acagcatgcg gcccgtggtc ctcctaagtg tgagcttgcg gcggaccgag gcccacctgc ctccctgcct gcttcgcca ggactcgtga ctgcgtccgc agaagaaatc acaacagcgc tggaattgct agtttgctag gcagcatctt ttggacctgc gaaccatatg catttcacct caaatctgtt tccaagttga aaacctttgg gtctttctat gcgaacggat tgaagaaacg caaaaagttt ctacggactt taaattaaa atg gaa aaa tat gaa aac ctg ggt ttg gtt gga Met Glu Lys Tyr Glu Asn Leu Gly Leu Val Gly 1 5 10</pre>	60 120 180 240 300 360 420 472

WO 99/31236

301

gaa Glu	ggg Gly	agt Ser	tat Tyr 15	gga Gly	atg Met	gtg Val	atg Met	aag Lys 20	tgt Cys	agg Arg	aat Asn	aaa Lys	gat Asp 25	act Thr	gga Gly	520
aga Arg	att Ile	gtg Val 30	gcc Ala	ata Ile	aag Lys	aag Lys	ttc Phe 35	tta Leu	gaa Glu	agt Ser	gac Asp	gat Asp 40	gac Asp	aaa Lys	atg Met	568
gtt Val	aaa Lys 45	aag Lys	att Ile	gca Ala	atg Met	cga Arg 50	gaa Glu	gtc Val	aag Lys	tta Leu	cta Leu 55	aag Lys	caa Gln	ctt Leu	agg Arg	616
cat His 60	gaa Glu	aac Asn	ttg Leu	gtg Val	aat Asn 65	ctc Leu	ttg Leu	gaa Glu	gtg Val	tgt Cys 70	aaa Lys	aaa Lys	aaa Lys	a		659
<21 <21	0> 12 1> 30 2> Di 3> Ho	01	sapie	ens												
	l> CI	os 328	33		,											
<22	2 > 38 3 > Vo so	ig_pe 885 on He core eq Li	5 ≥ijne 4.1	e mat	rix PVLS/	'TL										
		olyA_ 572		nal												
	0> 12 ctgaa		ccag	gaaco	cc to	aatg	gaggt	t ctt	caag			s Ar			g cca 1 Pro	55
gct Ala -10	acc Thr	agc Ser	ctg Leu	gct Ala	ggc Gly -5	cct Pro	gtc Val	ctg Leu	tcc Ser	acc Thr 1	ctc	att	gcc Ala	cca Pro 5	act Thr	103
CCC	atg Met	ttg Leu	ttt Phe 10	tgt Cys	gaa	gat Asp	aaa Lys	agc Ser 15	tgg Trp	gat Asp	ctt Leu	ttt Phe	ctt Leu 20	ttt Phe	ttt Phe	151
aag Lys	tct Ser	cac His 25	aag	aca Thr	tgg Trp	ggc Gly	atc Ile 30	tcc Ser	aca Thr	aat Asn	tta Leu	agt Ser 35	tcc Ser	tgt Cys	cca Pro	199
ttt Phe	gga Gly 40	aat	ttg Leu	ttt Phe	cta Leu	tgt Cys 45	gta Val	cag Gln	ttt Phe	gtc Val	aga Arg 50	gaa Glu	aaa Lys	caa Gln	agt Ser	247

ttt tgt atg aat aca gaa tgt gat tta cgc aag aat tgacaaaaaa Phe Cys Met Asn Thr Glu Cys Asp Leu Arg Lys Asn

<210> 128

aaaaaaa

55

<211> 477

<212> DNA

<213> Homo sapiens

<221> CDS <222> 121477	
<pre>&lt;221&gt; sig_peptide &lt;222&gt; 121288 &lt;223&gt; Von Heijne matrix     score 3.5</pre>	
seq SSCADSFVSSSSS/QP	
<400> 128 cctcggagca ggcggagtaa agggacttga gcgagccagt tgccggatta ttctatttcc	60
cetecetete teeegeeeg tatetettt caccettete ceacectege tegegtagee	120 168
atg gcg gag ccg tcg gcg gcc act cag tcc cat tcc atc tcc tcg tcg  Met Ala Glu Pro Ser Ala Ala Thr Gln Ser His Ser Ile Ser Ser  -55 -50 -45	168
tcc ttc gga gcc gag ccg tcc gcg ccc ggc ggc ggc ggg agc cca gga Ser Phe Gly Ala Glu Pro Ser Ala Pro Gly Gly Gly Gly Ser Pro Gly -40 -35 -30 -25	216
gcc tgc ccc gcc ctg ggg acg aag agc tgc agc tcc tcc tgt gcg gat Ala Cys Pro Ala Leu Gly Thr Lys Ser Cys Ser Ser Ser Cys Ala Asp	264
-20 -15 -10  tcc ttt gtt tct tcc tct tcc tct cag cct gta tct cta ttt tcg acc  Ser Phe Val Ser Ser Ser Ser Gln Pro Val Ser Leu Phe Ser Thr	312
-5 1 5 tca caa gag gga ttg agc tct ctt tgc tct gat gag cca tct tca gaa Ser Gln Glu Gly Leu Ser Ser Leu Cys Ser Asp Glu Pro Ser Ser Glu	360
10 15 20 att atg act tot toc ttt ott toa tot tot gaa ata cat aac act ggc	408
Ile Met Thr Ser Ser Phe Leu Ser Ser Glu Ile His Asn Thr Gly 25 30 35 40	
ctt aca ata cta cat gga gaa aaa agc cat gtg tta ggg agc cag cct Leu Thr Ile Leu His Gly Glu Lys Ser His Val Leu Gly Ser Gln Pro 45 50 55	456
att tta gcc aaa aaa aaa Ile Leu Ala Lys Lys Lys 60	477
•	
<210> 129 <211> 323 <212> DNA <213> Homo sapiens	
<220>	
<221> CDS <222> 2163	
<221> polyA_signal <222> 292297	
<221> polyA_site <222> 310323	
<pre>&lt;400&gt; 129 a gct ttc gtg tgg gag cca gct atg gtg cgg atc aat gcg ctg aca gca Ala Phe Val Trp Glu Pro Ala Met Val Arg Ile Asn Ala Leu Thr Ala 1 5 10 15</pre>	49
gcc tct gag gct gcg tgc ctg atc gtg tct gta gat gaa acc atc aag Ala Ser Glu Ala Ala Cys Leu Ile Val Ser Val Asp Glu Thr Ile Lys	97
20 25 30 aac ccc cgc tcg act gtg gat gct ccc aca gca gca ggc cgg ggc cgt	145

			cgc			tgag	aggc	ac c	ccac	ccat	c ac	atgg	ctgg			3
-	50		Arg													
ctaa	ctac	tg g	gtgc	actt	a cc	ctcc	ttgg	ctt	ggtt	act	tcat	ttta	ca a	ggaa	ggggt	3
agta	attg	gc c	cact	ctct	t ct	tact	ggag	gct	attt	aaa	taaa	atgt	aa g	actt	caaaa	3
aaaa	aaaa	aa														•
<210	> 13	0														
	.> 13															
	> DN															
<213	> HC	omo s	apie	ns												
<220	)>															
<221	.> CI	S														
<222	> 46	67	5													
	_ •			_												
	.> 51 ?> 46		ptid	.e												
			ijne	mat	rix											
		ore	-													
	se	eq LI	LLGL	SFIL	AGL/	IV										
				- 7												
	_		_sign _1369													
< 4 4 4	. > 13		1303													
<221	l> pc	olyA_	site	<b>:</b>												
	_		site 1392													
<222	2 > 13	883.	-													
<222	2 > 13 0 > 13	383 30	1392	!	ra aa	aaqa	aggg	c too	etctg	igga	gato	jt at	g ct	it ac	ct ctc	
<222 <400 ctco	2> 13 0> 13 cgagt	883 30 Etg (	1392 ccacc	cagg								Me	et Le	eu Ti	ct ctc nr Leu	
<222 <400 ctcc	2 > 13 0 > 13 cgagt	883	1392 ccacc	cagg	atc	ttg	qca	gga	ctt	att	gtt	ggt	et Le gga	eu Ti gcc	nr Leu tgc	
<222 <400 ctco tta Leu	2 > 13 0 > 13 cgagt	883	1392 ccacc	cagg	atc Ile	ttg	qca		ctt	att Ile	gtt	ggt	et Le gga	gcc Ala	nr Leu tgc	
<222 <400 ctcc tta Leu -10	2> 13 0> 13 cgagt ggc Gly	sesso etg o	1392 ccacc tca Ser	cagg ttc Phe	atc Ile -5	ttg Leu	gca Ala	gga Gly	ctt Leu	att Ile 1 .	gtt Val	ggt Gly	gga Gly	gcc Ala 5	tgc Cys	
<222 <400 ctco tta Leu -10 att	2 > 13 0 > 13 cgagt ggc Gly tac	stg of the Leu	tca Ser	cagg ttc Phe ttc	atc Ile -5 atq	ttg Leu ccc	gca Ala aag	gga Gly agc	ctt Leu acc	att Ile 1 .	gtt Val tac	ggt Gly cgt	gga Gly gga	gcc Ala 5 gag	tgc Cys atg	
<222 <400 ctco tta Leu -10 att	2 > 13 0 > 13 cgagt ggc Gly tac	stg of the Leu	tca ser tac	cagg ttc Phe ttc	atc Ile -5 atq	ttg Leu ccc	gca Ala aag	gga Gly	ctt Leu acc	att Ile 1 .	gtt Val tac	ggt Gly cgt	gga Gly gga	gcc Ala 5 gag	tgc Cys atg	
<222 <400 ctco tta Leu -10 att Ile	2> 13 0> 13 cgagt ggc Gly tac Tyr	stg of the control of	tca Ser tac Tyr	ttc Phe ttc Phe	atc Ile -5 atg Met	ttg Leu ccc Pro	gca Ala aag Lys cct	gga Gly agc ser 15 gca	ctt Leu acc Thr	att Ile 1 . att Ile	gtt Val tac Tyr	ggt Gly cgt Arg	gga Gly gga Gly 20 gga	gcc Ala 5 gag Glu	tgc Cys atg Met	
<222 <400 ctco tta Leu -10 att Ile	2> 13 0> 13 cgagt ggc Gly tac Tyr	stg of the control of	tca Ser tac Tyr	ttc Phe ttc Phe	atc Ile -5 atg Met	ttg Leu ccc Pro	gca Ala aag Lys cct Pro	gga Gly agc Ser	ctt Leu acc Thr	att Ile 1 . att Ile	gtt Val tac Tyr	ggt Gly cgt Arg cgt	gga Gly gga Gly 20 gga	gcc Ala 5 gag Glu	tgc Cys atg Met	
<222 <400 ctcc tta Leu -10 att Ile tgc Cys	2> 13 0> 13 cgagt ggc Gly tac Tyr ttt Phe	aag Lys ttt Phe	tca Ser tac Tyr 10 gat Asp	ttc Phe ttc Phe tct	atc Ile -5 atg Met gag Glu	ttg Leu ccc Pro gat Asp	gca Ala aag Lys cct Pro 30	gga Gly agc Ser 15 gca Ala	ctt Leu acc Thr aat Asn	att Ile 1 . att Ile tcc Ser	gtt Val tac Tyr ctt Leu	ggt Gly cgt Arg cgt Arg	gga Gly gga Gly 20 gga Gly	gcc Ala 5 gag Glu gga Gly	tgc Cys atg Met gag Glu	
<222 <400 ctco tta Leu -10 att Ile tgc Cys	2> 13 0> 13 cgagt ggc Gly tac Tyr ttt Phe	aag Lys ttt Phe 25	tca Ser tac Tyr 10 gat Asp	ttc Phe ttc Phe tct Ser	atc Ile -5 atg Met gag Glu	ttg Leu ccc Pro gat Asp	gca Ala aag Lys cct Pro 30 gag	gga Gly agc Ser 15 gca Ala	ctt Leu acc Thr aat Asn	att Ile 1 . att Ile tcc Ser	gtt Val tac Tyr ctt Leu	ggt Gly cgt Arg cgt Arg	gga Gly gga Gly 20 gga Gly	gcc Ala 5 gag Glu gga Gly	tgc Cys atg Met gag Glu	
<222 <400 ctco tta Leu -10 att Ile tgc Cys	2> 13 D> 13 Egagt ggc Gly tac Tyr ttt Phe	aag Lys ttt Phe 25	tca Ser tac Tyr 10 gat Asp	ttc Phe ttc Phe tct Ser	atc Ile -5 atg Met gag Glu	ttg Leu ccc Pro gat Asp act	gca Ala aag Lys cct Pro 30 gag Glu	gga Gly agc Ser 15 gca Ala	ctt Leu acc Thr aat Asn	att Ile 1 . att Ile tcc Ser	gtt Val tac Tyr ctt Leu att Ile	ggt Gly cgt Arg cgt Arg	gga Gly gga Gly 20 gga Gly	gcc Ala 5 gag Glu gga Gly	tgc Cys atg Met gag Glu	
<222 <400 ctco tta Leu -10 att Ile tgc Cys cct Pro	2> 13 cgagt ggc Gly tac Tyr ttt Phe aac Asn 40	aag Lys ttt Phe 25 ttc	tca Ser tac Tyr 10 gat Asp	ttc Phe ttc Phe tct Ser cct	atc Ile -5 atg Met gag Glu gtg Val	ttg Leu ccc Pro gat Asp act Thr	gca Ala aag Lys cct Pro 30 gag Glu	gga Gly agc Ser 15 gca Ala gag Glu	ctt Leu acc Thr aat Asn gct Ala	att Ile 1 . att Ile tcc Ser gac Asp	gtt Val tac Tyr ctt Leu att Ile 50	ggt Gly cgt Arg cgt Arg 35 cgt Arg	gga Gly gga Gly 20 gga Gly gag Glu	gcc Ala 5 gag Glu gga Gly gat Asp	tgc Cys atg Met gag Glu gac Asp	
<222 <400 ctco tta Leu -10 att Ile tgc Cys cct Pro	2> 13 cgagt ggc Gly tac Tyr ttt Phe aac Asn 40 att	aag Lys ttt Phe 25 ttc Phe	tca Ser tac Tyr 10 gat Asp ctg Leu	ttc Phe ttc Phe tct Ser cct Pro	atc Ile -5 atg Met gag Glu gtg Val	ttg Leu ccc Pro gat Asp act Thr 45 gtg	gca Ala aag Lys cct Pro 30 gag Glu	gga Gly agc Ser 15 gca Ala gag Glu	ctt Leu acc Thr aat Asn gct Ala	att Ile 1 . att Ile tcc Ser gac Asp	gtt Val tac Tyr ctt Leu att Ile 50	ggt Gly cgt Arg cgt Arg 35 cgt Arg	gga Gly gga Gly 20 gga Gly gag Glu	gcc Ala 5 gag Glu gga Gly gat Asp	tgc Cys atg Met gag Glu gac Asp	
<222 <400 ctcc tta Leu -10 att Ile tgc Cys cct Pro aac Asn 55	2> 13 cgagt ggc Gly tac Tyr ttt Phe aac Asn 40 att Ile	aag Lys ttt Phe 25 ttc Phe gca	tca Ser tac Tyr 10 gat Asp ctg Leu atc	ttc Phe ttc Phe tct Ser cct Pro	atc Ile -5 atg Met gag Glu gtg Val gat Asp	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val	gca Ala aag Lys cct Pro 30 gag Glu cct Pro	gga Gly agc Ser 15 gca Ala gag Glu gtc Val	ctt Leu acc Thr aat Asn gct Ala ccc Pro	att Ile 1 . att Ile tcc Ser gac Asp agt Ser 65	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe	ggt ggt ggt arg cgt Arg 35 cgt Arg	gga Gly gga Gly 20 gga Gly gag Glu gat Asp	gcc Ala 5 gag Glu gga Gly gat Asp agt	tgc Cys atg Met gag Glu gac Asp gac Asp	
<222 <400 ctco tta Leu -10 att Ile tgc Cys cct Pro aac Asn 55 cct	2> 13 cgagt cgagt cgagt cgagt tac Tyr ttt Phe aac Asn 40 att Ile	aag Lys ttt Phe 25 ttc Phe Ala	tca Ser tac Tyr 10 gat Asp ctg Leu atc	ttc Phe ttc Phe tct Ser cct Pro att Ile	atc Ile -5 atg Met gag Glu gtg Val gat Asp 60 cat	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val	gca Ala aag Lys cct Pro 30 gag Glu cct Pro	gga Gly agc Ser 15 gca Ala gag Glu gtc Val	ctt Leu acc Thr aat Asn gct Ala ccc Pro	att Ile 1 . att Ile tcc Ser gac Asp agt Ser 65 gga	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe	ggt ggt arg cgt arg set arg tct ser act	gga Gly gga Gly 20 gga Gly gag Glu gat Asp	gcc Ala 5 gag Glu gga Gly gat Asp agt tac	tgc Cys atg Met gag Glu gac Asp gac Asp	
<222 <400 ctco tta Leu -10 att Ile tgc Cys cct Pro aac Asn 55 cct	2> 13 cgagt cgagt cgagt cgagt tac Tyr ttt Phe aac Asn 40 att Ile	aag Lys ttt Phe 25 ttc Phe Ala	tca Ser tac Tyr 10 gat Asp ctg Leu atc	ttc Phe ttc Phe tct cct Pro att Ile	atc Ile -5 atg Met gag Glu gtg Val gat Asp 60 cat	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val	gca Ala aag Lys cct Pro 30 gag Glu cct Pro	gga Gly agc Ser 15 gca Ala gag Glu gtc Val	ctt Leu acc Thr aat Asn gct Ala ccc Pro aag Lys	att Ile 1 . att Ile tcc Ser gac Asp agt Ser 65 gga	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe	ggt ggt arg cgt arg set arg tct ser act	gga Gly gga Gly 20 gga Gly gag Glu gat Asp	gcc Ala 5 gag Glu gga Gly gat Asp agt tac Tyr	tgc Cys atg Met gag Glu gac Asp gac Asp	
<222 <400 ctco tta Leu -10 att Ile tgc Cys cct Pro aac Asn 55 cct	2> 13 cgagt ggc Gly tac Tyr ttt Phe aac Asn 40 att Ile gca Ala	aag Lys ttt Phe 25 ttc Phe gca Ala gca Ala	tca Ser tac Tyr 10 gat Asp ctg Leu atc Ile	ttc Phe ttc Phe tct Ser cct Pro att Ile att Ile	atc Ile -5 atg Met gag Glu gtg Val gat Asp 60 cat His	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val gac Asp	gca Ala aag Lys cct Pro 30 gag Glu cct Pro ttt	gga Gly agc Ser 15 gca Ala gag Glu gtc Val gaa Glu	ctt Leu acc Thr aat Asn gct Ala ccc Pro aag Lys	att Ile 1 . att Ile tcc Ser gac Asp agt Ser 65 gga Gly	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe atg Met	ggt ggt ggt cgt Arg cgt Arg tct Ser act	gga Gly gga Gly 20 gga Gly gag Glu gat Asp	gcc Ala 5 gag Glu ggay gat Asp agt tac Tyr 85	tgc Cys atg Met gag Glu gac Asp gac Asp 70 ctg Leu	
<222 <400 ctco tta Leu -10 att Ile tgc Cys cct Pro aac Asn 55 cpro gac	2> 13 cgagt cgagt ggc Gly tac Tyr ttt Phe aac Asn 40 att Ile gca Ala	aag ttt Leu aag Lys ttte 25 ttc Phe gca Ala ttg	tca Ser tac Tyr 10 gat Asp ctg Leu atc Ile	ttc Phe ttc Phe tct Ser cct Pro att Ile att Ile	atc Ile -5 atg Met gaglu gtg Val gat Asp 60 this atc	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val gac Asp	gca Ala aag Lys cct Pro 30 gag Glu cct Pro ttt Phe	gga Gly agc Ser 15 gca Ala gag Glu gtc Val gaa Glu ctg	ctt Leu acc Thr aat Asn gct Ala ccc Pro aag Lys 80 atg	att Ile 1 . att Ile tcc Ser gac Asp agt Ser Gly ccc	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe atg Met	ggt ggt arg cgt arg tct ser act Thr aat	gga Gly gga Gly 20 gga Gly gag Glu gat Asp	gcc Ala 5 gag Glu ggay gat Asp agt tac Tyr 85 tct	tgc Cys atg Met gag Glu gac Asp gac Asp 70 ctg Leu	
<222 <400 ctco tta Leu -10 att Ile tgc Cys cct Pro aac Asn 55 cpro gac	2> 13 cgagt cgagt ggc Gly tac Tyr ttt Phe aac Asn 40 att Ile gca Ala	aag ttt Leu aag Lys ttte 25 ttc Phe gca Ala ttg	tca Ser tac Tyr 10 gat Asp ctg Leu atc Ile att Ile	ttc Phe ttc Phe tct Ser cct Pro att Ile att Ile	atc Ile -5 atg Met gaglu gtg Val gat Asp 60 this atc	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val gac Asp	gca Ala aag Lys cct Pro 30 gag Glu cct Pro ttt Phe	gga Gly agc Ser 15 gca Ala gag Glu gtc Val gaa Glu	ctt Leu acc Thr aat Asn gct Ala ccc Pro aag Lys 80 atg	att Ile 1 . att Ile tcc Ser gac Asp agt Ser Gly ccc	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe atg Met	ggt ggt arg cgt arg tct ser act Thr aat	gga Gly gga Gly 20 gga Gly gag Glu gat Asp	gcc Ala 5 gag Glu ggay gat Asp agt tac Tyr 85 tct	tgc Cys atg Met gag Glu gac Asp gac Asp 70 ctg Leu	
<pre>&lt;222 &lt;400 ctcd ttau-10 atttle tCys ccto aasn 55cto gasp gtt</pre>	2> 13 0> 13 0> 13 0 gagt ggc Gly tac Tyr ttt Phe aac Asn 40 att Ile gca Ala ttgu	and control of the co	tcacce tca Ser tac Tyr 10 gat Asp ctg Leu atc Ile ctg Leu ctg Leu ctg Ceu cca	ttc Phe ttc Phe tct Ser cct Pro att Ile att Ile 75 ggly aaa	atc Ile -5 atg Met gaglu gtgl Val gat Asp 60 this atc Ile aat	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val gac Asp tgc Cys	gca Ala aag Lys cct Pro 30 gag Glu cct Pro ttt Phe tat Tyr	gga Gly agc Ser 15 gca Ala gag Glu gtc Val gaa Glu ctg Leu 95 gag	ctt Leu acc Thr aat Asn gct Ala ccc Pro aag Lys 80 atg Met	att Ile 1 . att Ile tcc Ser gac Asp agt 65 ag Gly ccc Pro	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe atg Met ctc Leu	ggt ggt arg cgt arg tct act Thr aat aaa	gga Gly gga Gly 20 gga Gly gag Glu gat Asp gct Ala act 100 ctg	gCla SagGlu GGly GGly GASP ASET TYS TEST	tgc Cys atg Met gag Glu gac Asp gac Asp 70 ctg Leu att Ile	
<pre>&lt;222 &lt;400 ctcd ttau-10 atttle tCys ccto aasn 55cto gasp gtt</pre>	2> 13 0> 13 0> 13 0 gagt ggc Gly tac Tyr ttt Phe aac Asn 40 att Ile gca Ala ttgu	and control of the co	tcacce tca Ser tac Tyr 10 gat Asp ctg Leu atc Ile ctg Leu ctg Leu ctg Ceu cca	ttc Phe ttc Phe tct Ser cct Pro att Ile att Ile 75 ggly aaa	atc Ile -5 atg Met gaglu gtgl Val gat Asp 60 this atc Ile aat	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val gac Asp tgc Cys	gca Ala aag Lys cct Pro 30 gag Glu cct Pro ttt Phe tat Tyr	gga Gly agc Ser 15 gca Ala gag Glu gtc Val gaa Glu ctg Leu 95 gag	ctt Leu acc Thr aat Asn gct Ala ccc Pro aag Lys 80 atg Met	att Ile 1 . att Ile tcc Ser gac Asp agt 65 ag Gly ccc Pro	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe atg Met ctc Leu	ggt ggt arg cgt arg tct act Thr aat aaa	gga Gly gga Gly 20 gga Gly gag Glu gat Asp gct Ala act 100 ctg	gCla SagGlu GGly GGly GASP ASET TYS TEST	tgc Cys atg Met gag Glu gac Asp gac Asp 70 ctg Leu att Ile	
<222 <400 ctc tta Leu -10 att le tgc cys cro aasn 55ct Pro gasp yal	2> 13 0> 13 0> 13 0 gagt 0 ggc Gly tac Tyr ttt Phe aac Asn 40 att Ile gca Ala tteu atg	and the second s	tca Ser tac Tyr 10 gat Asp ctg Leu atc Ile att Ile ctg Leu Pro	ttc Phe ttc Phe tct Ser cct Pro att Ile att Ile 75 gggy aaa Lys	atc Ile -5 atg Met gaglu gtgl Val gat Asp 60 this atc Ile aat Asn	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val gac Asp tgc Cys ctg	gca Ala aag Lys cct Pro 30 gag Glu cct Pro ttt Phe tat Tyr gta Val	gga Gly agc Ser 15 gca Ala gag Glu gtc Val gaa Glu ctg Leu 95 gag Glu	ctt Leu acc Thr aat Asn gct Ala ccc Pro aag Lys 80 atg Met ctc Leu	att Ile 1 . att Ile tcc Ser gac Asp agt Ser 65 a Gly ccc Pro ttt Phe	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe atg Met ctc Leu	ggt	gga Gly gga Gly 20 gga Gly gat Asp gct Ala act 100 ctg	gCC Ala 5 gag Glu ggay gatp ager tarr 85 ter gcal	tgc Cys atg Met gag Glu gac Asp gac Asp 70 ctg Leu att Ile agt Ser	
<222 <400 ctc ttau-10 atte -10 atte Cys cpr aasn 5cpr gasp yal	2> 13 0> 13 0> 13 0> 13 0> 13 0	and the control of th	tcacce tca Ser tac Tyr 10 gat Asp ctg Leu atc Ile att Ile ctgu cro ctg	ttc Phe ttc Phe tctr cro atte Ile atte 75 96ly aaa Lys	atcelle -5 get Met gagu gtal Val gatp 60 this atcelle aatn caa	ttg Leu ccc Pro gat Asp act Thr 45 gtg Val gac Asp tgc Cys ctg	gca Ala aag Lys cct Pro 30 gag Glu cct Pro ttt Phe tat Tyr gta Val 110 tat	gga Gly agc ser 15 gca Ala galu gtc Val gta Clu ctg Glu ctg Glu ctg Glu	ctt Leu acc Thr aat Asn gct Ala cco Pro adys atg Met cteu gtt	att Ile 1 .tt Ile tccr gasp ager 65 aggly cco tthe cga	gtt Val tac Tyr ctt Leu att Ile 50 ttc Phe atg Met cteu ggc Gly gaa	gty cgtg cgtg carg tcr actr actr actr actr actr actr actr	gga Gly ggay ggly gat Asp gct Act Throctg Leu cta	gCla gGlu gGly gAsp tarr street gAla gtt	tgc Cys atg Met gag Glu gac Asp gac Asp 70 ctg Leu att Ile agt Ser	

•	
gtg gag gaa att cgt gat gtt agt aac ctt ggc atc ttt att tac caa	537
Val Glu Glu Ile Arg Asp Val Ser Asn Leu Gly Ile Phe Ile Tyr Gin	
135 140 145 · 150	585
ctt tgc aat aac aga aag tcc ttc cgc ctt cgt cgc aga gac ctc ttg Leu Cys Asn Asn Arg Lys Ser Phe Arg Leu Arg Arg Arg Asp Leu Leu	303
Leu Cys Asn Asn Arg Lys Ser Phe Arg Leu Arg Arg Arg Asp 200 165	
ctg ggt ttc aac aaa cgt gcc att gat aaa tgc tgg aag att aga cac	633
Leu Gly Phe Asn Lys Arg Ala Ile Asp Lys Cys Trp Lys Ile Arg His	
170 175 180	
ttc ccc aac gaa ttt att gtt gag acc aag atc tgt caa gag	675
Phe Pro Asn Glu Phe Ile Val Glu Thr Lys Ile Cys Gln Glu	
185 190 195	735
taagaggcaa cagatagagt gtccttggta ataagaagtc agagatttac aatatgactt	795
taacattaag gtttatggga tactcaagat atttactcat gcatttactc tattgcttat	855
gctttaaaaa aaggaaaaaa aaaaaactac taaccactgc aagctcttgt caaattttag tttaattggc attgcttgtt ttttgaaact gaaattacat gagtttcatt ttttctttgc	915
atttataggg tttagatttc tgaaagcagc atgaatatat cacctaacat cctgacaata	975
aattocatoo gttgtttttt ttgtttgttt gttttttctt ttcctttaag taagetettt	1035
artcatctta tootooagca attttaaaaat ttqaaatatt ttaaattgtt tttgaacttt	1095
ttototaaaa tatatcagat ctcaacattg ttggtttctt ttgtttttca ttttgtacaa	1155
ctttcttgaa tttagaaatt acatctttgc agttctgtta ggtgctctgt aattaacctg	1215
acttatatot gaacaatttt catgagacag tcatttttaa ctaatgcagt gattctttct	1275
cactactate tgtattgtgg aatgcacaaa attgtgtagg tgctgaatge tgtaaggagt	1335
ttaggttgta tgaattctac aaccctataa taaattttac tctatacaaa aaaaaaa	1392
•	
<210> 131	
<211> 999	
<212> DNA	
<213> Homo sapiens	
· ·	
<220>	
<221> CDS	
<222> 62385	
.201. molyn gignal	
<221> polyA_signal	
(222) 9/49/J	
<221> polyA_site	
<222> 987999	
<400> 131	60
cotgaatgac ttgaatgttt coccgootga gotaacagto catgtgggtg attcagotot	109
g atg gga tgt gtt ttc cag agc aca gaa gac aaa tgt ata ttc aag ata	100
Met Gly Cys Val Phe Gln Ser Thr Glu Asp Lys Cys Ile Phe Lys Ile	3
gac tgg act ctg tca cca gga gag cac gcc aag gac gaa tat gtg cta	157
Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu	
20 25 30	
tac tat tac tcc aat ctc agt gtg cct att ggg cgc ttc cag aac cgc	205
Tyr Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg	
35 40 45	
gta cac ttg atg ggg gac atc tta tgc aat gat ggc tct ctc ctg ctc	253
Val His Leu Met Gly Asp Ile Leu Cys Asn Asp Gly Ser Leu Leu Leu	
50 55 60	301
caa gat gtg caa gag gct gac cag gga acc tat atc tgt gaa atc cgc	301
Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg 65 70 75 80	
70	
ctc ass one dag and can did tic asd asd ded did did cat did	349
ctc aaa ggg gag agc cag gtg ttc aag aag gcg gtg gta ctg cat gtg Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val	349
ctc aaa ggg gag agc cag gtg ttc aag aag gcg gtg gta ctg cat gtg Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val 85 90 95	349

• •	
ctt cca gag gag ccc aaa ggt acg caa atg ctt act taaagagggg	395
ctt cca gag gag ccc dad ggt aby cla day to the	
Leu Pro Glu Glu Pro Lys Gly Thr Gln Met Leu Thr	•
100 105	455
ccaaggggca agagctttca tgtgcaagag gcaaggaaac tgattatctt gagtaaatgc	
carcetting detaagtact taccacagag tgaatettea aaaaatgate ataattatte	515
cagtcaataa aaatagagtt attttattaa ataaaatatt gataattatt gtattattac	575
tttaaacaca cttcccctc acaaaagccc tgtgaaggat gttttgttca catatatgtc	635
tttaaacaca cttccccctc acaaaagccc tgtgaagaac goodaaa ccctgcacc	695
caaatatgtt ttggacacat atttattaaa tggaataaat agtacttgaa ccctggcacc	755
totgacaaca aagtocatgt tottttact atgocctaat acctttcatc agttatocac	815
attratorta catchoratt thataggiac cotatottag gigitologge ggalagada	
gaaataagga ggccaggctc agtggctcat gcctgtaatc ctagcatttt gggaggctga	875
ggcagcagaa ctgcctgagc cccagggttc aagactgcag tgagctatga tggcaccact	935
gcattctagc ctgggtgaca gagcaagact ctgtctaaaa taaaaaaaga gaaaaaaaa	995
gcattctage ctgggtgaca gagcaagace ctgtctada tadaman 5	999
aaaa	
<210> 132	
<211> 725	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 422550	
<222> 422350	
<221> sig_peptide	
<222> 422475	
<223> Von Heijne matrix	
score 4.5	
seq LRWLMPVIPALWG/AE	
and make aire	
<221> polyA_site	
<222> 714725	
<400> 132	60
tctgcgaggg tgggagagaa aattaggggg agaaaggaca gagagagcaa ctaccatcca	
taggragata ggtgagtaaa tatatttgca qtaacctatt tgctattcct tgctgcaact	120
gtgtttaatg ttccttccag aatcagagag agtattgcca tccaagaaat cgtttttaga	180
tatgacattt gagctatcat cttgagacca atacctaaaa caatttcagt ttaagaaatg	240
tatgacattt gagetateat ettgagacea attestaat gaatttatta ceetcagega	300
totaggtatg gtgaaaacac agtttaaaac cagcaaaaca gaatttattg ccctcagcga	360
atacccacaa tgtacatata ccttgtattt ctgaaagcaa agcaagcatg ccaagtagtt	420
tttatttacc tgtacctata atacagcaag gtgaaacagg atatatttit gaagttada	
a ato tot toa ogo ogo oto ogo too oto ato oot gta ato ood yea eet	469
Met Ser Ser Gly Arg Leu Arg Trp Leu Met Pro Val Ile Pro Ala Leu	
-15	517
tgg gga gcc gag aag ggt gaa tca cct gag gtc agc agt ttt gag acc	
Trp Gly Ala Glu Lys Gly Glu Ser Pro Glu Val Ser Ser Phe Glu Thr	
1 5 10	
agg ctg gcc aac atg gcg aaa ccc tgt ctc tac tgaaaataca aaaattagct	570
Arg Leu Ala Asn Met Ala Lys Pro Cys Leu Tyr	
	630
gggtgtggtg gcgggcgcct gtagtcccag ctacttggga gactgaggca ggagaattgc	690
ttgaacacgg aaggcggaag ttgcagtaag ctgagatcgt gccaccgcac accagcttgg	725
gcaacagagt gagactccct ctcaaaaaaa aaaaa	123

<211> 400

<212> DNA

<213> Homo sapiens

<220> <221> CDS <222> 124231	
<221> polyA_site <222> 387400	
<pre>&lt;400&gt; 133 ctcgcctctc ctggcttctg gtatgcacca gcaattcctg gcgttccttg gctcctagaa gcatcactcc tatcacatgg tcatcttcac cctgtgtgtc ttcacactac cctttctctg tgc atg tct gcc cga atc cct ttt tat aag gac acc agt cag att aga     Met Ser Ala Arg Ile Pro Phe Tyr Lys Asp Thr Ser Gln Ile Arg     1</pre>	60 120 168
tta ggg tct acc ata ata cct cat ttt aac tta atc acc ttt gta aag Leu Gly Ser Thr Ile Ile Pro His Phe Asn Leu Ile Thr Phe Val Lys 20 25 30	216
acc ttt ttc caa ata tagtcactct ctgaggtact gatggttagg atctcaacat Thr Phe Phe Gln Ile 35	271
accttttttg ggaggacaca attgaaccca taacagggtg tttgcaagga agagttaaaa tttgaaagaa aggtggtatt tgcttagata gatagggcac agctttctag gtgacaaaaa aaaaaaaaa	331 391 400
<210> 134 <211> 1053 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 1311051	
<221> sig_peptide <222> 131169 <223> Von Heijne matrix score 4.2 seq MLAVSLTVPLLGA/MM	
<221> polyA_signal	
<pre>&lt;400&gt; 134 gagcgaggcg gacgggctgc gacagcgccg gcccctgcgg ccgcaggtcg tcacagacga tgatggccag gccccggagg ctaaggacgg cagctccttt agcggcagag ttttccgagt gaccttcttg atg ctg gct gtt tct ctc acc gtt ccc ctg ctt gga gcc</pre>	60 120 169
atg atg ctg ctg gaa tct cct ata gat cca cag cct ctc agc ttc aaa Met Met Leu Leu Glu Ser Pro Ile Asp Pro Gln Pro Leu Ser Phe Lys 1 10 15	217
gaa ccc ccg ctc ttg ctt ggt gtt ctg cat cca aat acg aag ctg cga Glu Pro Pro Leu Leu Gly Val Leu His Pro Asn Thr Lys Leu Arg 20 25 30	265
cag gca gaa agg ctg ttt gaa aat caa ctt gtt gga ccg gag tcc ata Gln Ala Glu Arg Leu Phe Glu Asn Gln Leu Val Gly Pro Glu Ser Ile 35 40 45	313
gca cat att ggg gat gtg atg ttt act ggg aca gca gat ggc cgg gtc Ala His Ile Gly Asp Val Met Phe Thr Gly Thr Ala Asp Gly Arg Val 50 55 60	361

gta Val	aaa Lys	ctt Leu	gaa Glu	aat Asn	Gly	gaa Glu	ata Ile	gag Glu	acc Thr	IIe	gcc Ala	cgg Arg	ttt Phe	ggt Gly	tcg Ser 80	409
65 ggc Gly	cct Pro	tgc Cys	aaa Lys	Thr	70 cga Arg	gat Asp	gat Asp	gag Glu	Pro	75 gtg Val	tgt Cys	999 999	aga Arg	ccc Pro 95	ctg	457
aat	atc	cat	gca Ala	85 aaa	ccc	aat	aaa	act Thr	90 ctc	ttt	gtg	gcc	gat	gca	tgc	505
aag Lys	gga Gly	Leu	100 ttt Phe	gaa Glu	gta Val	aat Asn	ccc Pro	105 tgg Trp	aaa Lys	cgt Arg	gaa Glu	gtg Val 125	aaa	ctg Leu	ctg Leu	553
ctg Leu	Ser	115 tcc Ser	gag Glu	aca Thr	ccc Pro	Ile	qaq	gly aaa	aag Lys	aac Asn	atg Met 140	tcc	ttt Phe	gtg Val	aat Asn	601
gat Asp	130 ctt Leu	aca Thr	gtc Val	tct Ser	Gln	135 gat Asp	ggg Gly	agg Arg	aag Lys	lle	tat	ttc Phe	acc Thr	gat Asp	tct Ser 160	649
145 agc Ser	agc Ser	aaa Lys	tgg Trp	Gln	150 aga Arg	cga Arg	gac Asp	tac Tyr	Leu	155 ctt Leu	ctg Leu	gtg Val	atg Met	gag Glu 175	ggc	697
aca Thr	gat Asp	gac Asp	gly ggg	165 cgc Arg	ctg Leu	ctg Leu	gag Glu	tat Tyr	170 gat Asp	act Thr	gtg Val	acc Thr	Arg	gaa	gta Val	745
aaa Lys	gtt Val	tta Leu	180 ttg Leu	gac Asp	cag Gln	ctg Leu	cgg Arg	185 ttc Phe	ccg Pro	aat Asn	gga Gly	Val	190 cag Gln	ctg Leu	tct Ser	793
cct	aca	195	gac	ttt	atc	cta	200 ata	gca Ala	qaa	aca	acc	205 atg	gcc	agg	ata	841
caa	210 aga	gtc	tac	att	tct	215 ggc	ctg	atg Met	aag	ggc	220 999	gct	gat	ctg	ttt Phe	889
225	aaa	aac	ato	cat	230 gga	ttt	cca	gac Asp	aac	235 atc	cgg	ccc	agc	agc Ser	tct	937
aaa	aaa	tac	taa	245 ata	aac	atq	tcg	acc Thr	250 atc	cgc	cct	aac	cct Pro	255 999	ttt	985
+cc	ato	cto	260	tto	tta	tct	gag	265 aqa	ccc	tgg	att	aaa Lys	agg Arg	atg		1033
ttt	. aag	275 gca Ala		aaa	aaa	aa	280	1				285				1053

<211> 1128 <212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 86..403

<221> sig_peptide

<222> 86..181

<223> Von Heijne matrix score 8.8

seq VPMLLLIVGGSFG/LR

```
<221> polyA_signal
<222> 1097..1102
<221> polyA_site
<222> 1117..1128
<400> 135
cgtcttggtg agagcgtgag ctgctgagat ttgggagtct gcgctaggcc cgcttggagt
                                                                       60
tctgagccga tggaagagtt cactc atg ttt gca ccc gcg gtg atg cgt gct
                                                                      112
                            Met Phe Ala Pro Ala Val Met Arg Ala
                                    -30
ttt cgc aag aac aag act ctc ggc tat gga gtc ccc atg ttg ttg ctg
                                                                      160
Phe Arg Lys Asn Lys Thr Leu Gly Tyr Gly Val Pro Met Leu Leu Leu
                                -15
            -20
att gtt gga ggt tot ttt ggt ott ogt gag ttt tot caa atc oga tat
Ile Val Gly Sly Ser Phe Gly Leu Arg Glu Phe Ser Gln Ile Arg Tyr
        -5
                                                                      256
gat gct gtg aag agt aaa atg gat cct gag ctt gaa aaa aaa ctg aaa
Asp Ala Val Lys Ser Lys Met Asp Pro Glu Leu Glu Lys Lys Leu Lys
                                        20
                    15
gag aat aaa ata tot tta gag tog gaa tat gag aaa ato aaa gac too
                                                                      304
Glu Asn Lys Ile Ser Leu Glu Ser Glu Tyr Glu Lys Ile Lys Asp Ser
                30
aag ttt gat gac tgg aag aat att cga gga ccc agg cct tgg gaa gat
                                                                      352
Lys Phe Asp Asp Trp Lys Asn Ile Arg Gly Pro Arg Pro Trp Glu Asp
                                50
            45
cet gae etc etc caa gga aga aat eca gaa age ett aag act aag aca
                                                                      400
Pro Asp Leu Leu Gln Gly Arg Asn Pro Glu Ser Leu Lys Thr Lys Thr
                                                 70
                             65
                                                                      453
act tgactctgct gattcttttt tccnnntttt ttttttttta aataaaaata
ctattaactg gacttcctaa tatatacttc tatcaagtgg aaaggaaatt ccaggcccat
ggaaacttgg atatgggtaa tttgatgaca aataatcttc actaaaggtc atgtacaggt
                                                                      573
ttttatactt cccagctatt ccatctgtgg atgaaagtaa caatgttggc cacgtatatt
                                                                      633
ttacacctcg aaataaaaaa tgtgaatact gctccaaaaa aaaaaaccag taccgtgtag
                                                                      693
tetetetegt ggettggatt tacactggge aacgtggttg gaatgtatet ggeteagaac
                                                                      753
tatgatatac caaacctggc taaaaaactt gaagaaatta aaaaggactt ggatgccaag
                                                                      813
                                                                      873
aagaaacccc ctagtgcatg agactgcctc cagcactgcc ttcaggatat accgattcta
ctgctcttga gggcctcgtt tactatctga accaaaagct tttgttttcg tctccagcct
                                                                      933
cagcacttet ettetteget agaccetgtg ttttttgett taaagcaage aaaatgggge
                                                                      993
cccaatttga gaactacccg acgtttccaa catactcacc tcttcccata atccctttcc
                                                                     1053
 aactgcatgg gaggttctaa gactggaatt atggtgctag attagtaaac atgactttta
                                                                     1113
                                                                      1128
 acqaaaaaa aaaaa
```

<211> 254

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 37..162

<221> sig_peptide

<222> 37..93

<223> Von Heijne matrix
score 9.5
seep-lmCLSLCTAFALS/KP

<221> polyA_signal <222> 224229	٠.
<221> polyA_site <222> 243254	
<400> 136 tgtgctgtgg gggctacgag gaaagatcta attatc atg gac ctg cga cag ttt Met Asp Leu Arg Gln Phe -15	54
ctt atg tgc ctg tcc ctg tgc aca gcc ttt gcc ttg agc aaa ccc aca Leu Met Cys Leu Ser Leu Cys Thr Ala Phe Ala Leu Ser Lys Pro Thr -10 -5 1	102
gaa aag aag gac cgt gta cat cat gag cct cag ctc agt gac aag gtt Glu Lys Lys Asp Arg Val His His Glu Pro Gln Leu Ser Asp Lys Val 5 10 15	150
cac aat gat att tgatagaacc aattgttgta cataaaacag atctgcgcat His Asn Asp Ile 20	202
atatatatat gtataaaaaa taataaaata atggaagatg aaaaaaaa	254
<210> 137 <211> 886 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 31381	
<221> sig_peptide	
<pre>&lt;222&gt; 3190 &lt;223&gt; Von Heijne matrix     score 5.4     seq AFVIACVLSLIST/IY</pre>	
<221> polyA_site <222> 875886	
<pre>&lt;400&gt; 137 ggaggatggg cgagcagtct gaatggcaga atg gat aac cgt ttt gct aca gca</pre>	54
ttt gta att gct tgt gtg ctt agc ctc att tcc acc atc tac atg gca Phe Val Ile Ala Cys Val Leu Ser Leu Ile Ser Thr Ile Tyr Met Ala -10 -5 1	102
gcc tcc att ggc aca gac ttc tgg tat gaa tat cga agt cca gtt caa Ala Ser Ile Gly Thr Asp Phe Trp Tyr Glu Tyr Arg Ser Pro Val Gln 5 10 15 20	150
gaa aat too agt gat ttg aat aaa ago ato tgg gat gaa tto att agt Glu Asn Ser Ser Asp Leu Asn Lys Ser Ile Trp Asp Glu Phe Ile Ser 25 30 35	198
gat gag gca gat gaa aag act tat aat gat gca ctt ttt cga tac aat Asp Glu Ala Asp Glu Lys Thr Tyr Asn Asp Ala Leu Phe Arg Tyr Asn 40 45 50	246
ggc aca gtg gga ttg tgg gga cgg tgt atc acc ata ccc aaa aac atg Gly Thr Val Gly Leu Trp Gly Arg Cys Ile Thr Ile Pro Lys Asn Met	294
cat tgg tat agc cca cca gaa agg aca ggt att tct ctt att tta act His Trp Tyr Ser Pro Pro Glu Arg Thr Gly Ile Ser Leu Ile Leu Thr	342

70 75 80	
tot gto tto tto acc tgg tta ata ata gac aaa acg acg taatgattgc Ser Val Phe Phe Thr Trp Leu Ile Ile Asp Lys Thr Thr	391
85 90 95 ccaattacat gtaagcaggt ttgttggttc tctctctctt taaagaaata aatcgtgtat	451
cttetette tactgeette tetecceaac ttetttgeat taccatggta etcateaata	511
tiggtiggat gaggaactit ictiatotig ggaaagcott aatggottit tittittotta	571
tttactcact cattaaaata cttttcatta ctctaacaca tgttataaag aaatagttgg	631
aaaagtgcat cgaaagactt ttaaaaatat ttggtaacta gtaaaaggac taccatcgaa	691 751
aatcaactca aaaaattgtc cttttatggg ttagctgtat tataatacat atctatcatt tgcccctgtg tcttagagga tataatttga ccagctctac atttaatctg tgtaattatg	811
agactgtttt acaacaatct tgatgcagag ttggtaggtt aagaaatttg tattacagaa	871
gttaaaaaaa aaaaa	886
<b>9</b>	
<210> 138	
<211> 1244 <212> DNA	
<213> Homo sapiens	
•	
<220>	
<221> CDS <222> 46579	
<2222 40373	
<221> sig_peptide	
<222> 46156	
<223> Von Heijne matrix	
score 3.5	
seq LVFNFLLILTILT/IW	
<400> 138	
cccttatcca ggttnttatc tanggaatcc cnnnaagact gggga atg gag aga cag	E 77
Met Glu Arg Gin	57
Met Glu Arg Gln -35	105
Met Glu Arg GIn -35 tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna	
Met Glu Arg Gln -35  tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa -30 -25 -20	
Met Glu Arg Gln -35  tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa -30 -25 -20  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg	
Met Glu Arg Gln  -35  tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  -25  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu	105
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  -10  Met Glu Arg Gln  -35	105
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  -10  -5  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act	105
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  -10  Met Glu Arg Gln  -35	105
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  Met Glu Arg Gln  -35  -35  -20  gcg cn cat cat can nna  -20  sca atc ttg ctt atc aat ttt ttg ctc atc ctt acc att ttg  -20  -20  sca atc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  1  1  1  1  1  1  1  1  1  1  1  1	105
Met Glu Arg Gln  -35  tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr	105 153 201
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20  Met Glu Arg Gln  -35  -35  -36  -20  5  -20  5  -20  5  -20  5  -10  -5  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  1  1  1  1  1  1  1  1  1  20  25  30	105 153 201 249
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20  gct aca gca cca act gat att gga agt gga act gtc tgt gac tgt gta	105 153 201
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30 -25 -20  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15 -10 -5  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1 5 10 15  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20 25 30  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt  Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val	105 153 201 249
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt gta  Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  35	105 153 201 249
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt gta  Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  35  40  45  aaa cta act gac caa	105 153 201 249 297
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30 -25 -20  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15 -10 -5  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1 5 10  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20 25 30  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt gta  Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  aaa cta act ttc agt cca cca act ctg ctg gtt aat gtc act gac caa  Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn Val Thr Asp Gln	105 153 201 249 297
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt gta  Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  35  aaa cta act ttc agt cca cca act ctg ctg gtt aat gtc act gac caa  Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn Val Thr Asp Gln  50  gtt tat gaa tat aaa tac aaa aga gaa ata agt cag cac aac atc aat	105 153 201 249 297
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  aaa cta act ttc agt cca cca act ctg ctg gtt aat gtc act gac caa Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn Val Thr Asp Gln  50  gtt tat gaa tat aaa tac aaa aga gaa ata agt cag cac aac atc aat Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln His Asn Ile Asn	105 153 201 249 297
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1	105 153 201 249 297 345
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15  aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt gta Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  35  aaa cta act ttc agt cca cca act ctg ctg gtt aat gtc act gac caa Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn Val Thr Asp Gln  50  gtt tat gaa tat aaa tac aaa aga gaa ata agt cag cac aac atc aat Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln His Asn Ile Asn  70  cct cat caa gga aat gtc ata ctt gaa aag atg aca ttt gat cca gaa	105 153 201 249 297
The color of the	105 153 201 249 297 345
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30 -25 -20  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15 -10 -5  aca atc tgg tta ttt aaa aat cat cgg ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1 5 10 15  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20 25  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt  Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  35 aaa cta act ttc agt cca cca act ctg ctg gtt aat gtc act gac caa  Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn Val Thr Asp Gln  50  gtt tat gaa tat aaa tac aaa aga gaa ata agt cag cac aac atc aat  Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln His Asn Ile Asn  65	105 153 201 249 297 345
Ser Arg Val   Met   Ser   Glu   Lys   Asp   Glu   Tyr   Gln   Phe   Gln   His   Xaa   Xaa   Xaa   Leu   Leu   Val   Phe   Asn   Phe   Leu   Leu   Thr   Ile   Leu   Leu   Thr   Ile   Trp   Leu   Phe   Lys   Asp   His   Arg   Phe   Arg   Phe   Leu   His   Glu   Thr   Ile   Glu   Thr   Ile   Glu   Thr   Ile   Trp   Leu   Leu   Val   Phe   Asn   Asn   Asn   Asn   Asn   Ala   Ile   Leu   Leu   Leu   Ile   Ser   Glu   Thr   Asp   Glu   Thr   Ile   Trp   Leu   Leu   Thr   Ile   Trp   Leu   Thr   Asp   Glu   Thr   Ile   Trp   Leu   Thr   Asp   Ile   Glu   Ser   Gly   Thr   Val   Cys   Asp   Cys   Val   Asn   Asn   Val   Tyr   Glu   Thr   Leu   Leu   Leu   Leu   Thr   Asp   Glu   Thr   Leu   Leu   Thr   Asp   Glu   Thr   Asp   Glu   Thr   Asp   Glu   Thr   Asp   Glu   Thr   Asp   Glu   Thr   Thr   Asp   Glu   Thr   Thr   Asp   Glu   Thr   Thr   Asp   Glu   Thr   Thr   Thr   Asp   Glu   Thr   Thr   Asp   Glu   Thr   Thr   Asp   Glu   Thr   Thr   Thr   Asp   Glu   Thr   Th	105 153 201 249 297 345 393
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna  Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa  -30 -25 -20  gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg  Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu  -15 -10 -5  aca atc tgg tta ttt aaa aat cat cgg ttc cgc ttc ttg cat gaa act  Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr  1 5 10 15  gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat  Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr  20 25  gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt  Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val  35 aaa cta act ttc agt cca cca act ctg ctg gtt aat gtc act gac caa  Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn Val Thr Asp Gln  50  gtt tat gaa tat aaa tac aaa aga gaa ata agt cag cac aac atc aat  Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln His Asn Ile Asn  65	105 153 201 249 297 345 393

agt cta aag aag aga cac ttt ttt caa aac tta gga tct att tta acg Ser Leu Lys Lys Arg His Phe Phe Gln Asn Leu Gly Ser Ile Leu Thr 115 120 125	537
tat gcc ttc ttg gga act gcc atc tcc tgc atc gtc ata ggg Tyr Ala Phe Leu Gly Thr Ala Ile Ser Cys Ile Val Ile Gly 130 135 140	579
taagtgacat teggagetea agttgeaggt ggetgtgggg tetgtgatet gtgtgaggga	639
totagoactt coaggattot tootgootgo gaaaattgto titititag talallacat	699 759
artrotator retrictoac traattocac qqcttctgac aaatacaagg cttcaaalca	819
aagcaaacta gaggattgct ggactttctc tgtgagttct ggacttctga cttagggaat gtggatcact tgccttgagt tatgtgaagc gcattgcatt	879
gegateagg teactgeatt ettetetete tegtategag agacettace tgtatetgge	939
aggagtgcaa aagtaactat atgccaaqaq ttttctttct aaaggaaagt ttacaagaca	999
grantiticaa aragataton tocaaatatn naacaqagtt gottaataca gggatagott	1059 1119
ttcagttaat accetgtaga atgeagaete tttnttteat tgtattttet tgattatget actgageeet aagteacaeg ttatataete tggettgeag eteateataa agtaaaatgt	1179
ggtaccaaat ggtgaaggca atccagcctn tgataatccc gtccaataca ttaaagntcc	1239
actgc	1244
<210> 139 <211> 471	
<212> DNA	
<213> Homo sapiens	
<220> <221> CDS	
<222> 92469	
<221> sig_peptide	
<222> 92172	
<pre>&lt;223&gt; Von Heijne matrix score 7.9</pre>	
seq VVVLALGFLGCYG/AK	
<221> polyA_signal <222> 454459	
<222> 454459	
<221> polyA_site	
<222> 458471	
400 100	
<pre>&lt;400&gt; 139 gcaagtgcag aagtcggtga cggtgggcat ctgggtgtca atcgatgggg catcctttct</pre>	60
gaagatette gggccactgt cgtccagtge c atg cag tit gic aac gig ggc	112
Met Gln Phe Val Asn Val Gly	
tac ttc ctc atc gca gcc ggc gtt gtg gtc ctt gct ctt ggt ttc ctg	160
Tyr Phe Leu Ile Ala Ala Gly Val Val Leu Ala Leu Gly Phe Leu	
-20 -15 -10 -5	
ggc tgc tat ggt gct aag act gag agc atg tgt gcc ctc gtg acg ttc	208
Gly Cys Tyr Gly Ala Lys Thr Glu Ser Met Cys Ala Leu Vai Thr Phe	
1 5 10 ttc ttc atc ctc ctc ctc atc ttc att gct gag gtt gca gct gct gtg	256
Phe Phe Ile Leu Leu Ile Phe Ile Ala Glu Val Ala Ala Val	
15 20 25	
gtc gcc ctg gtg tac acc aca atg gct gag cac ttc ctg acg ttg ctg	304
Val Ala Leu Val Tyr Thr Thr Met Ala Glu His Phe Leu Thr Leu Leu	
30 35 40 gta gtg cct gcc atc aag aaa gat tat ggt tcc cag gaa gac ttc act	352
Val Val Pro Ala Ile Lys Lys Asp Tyr Gly Ser Gln Glu Asp Phe Thr	
45 50 55 60	

the second secon	
caa gtg tgg aac acc acc atg aaa ggg ctc aag tgc cgt ggc ttc acc Gln Val Trp Asn Thr Thr Met Lys Gly Leu Lys Cys Arg Gly Phe Thr 65 70 75	400
aac tat acg gat ttt gag gac tca ccc tac ttc aaa atg cat aaa cct Asn Tyr Thr Asp Phe Glu Asp Ser Pro Tyr Phe Lys Met His Lys Pro 80 85 90	448
gtt aca atg aaa aaa aaa aa Val Thr Met Lys Lys Lys 95	471
<210> 140	
<211> 849	
<212> DNA <213> Homo sapiens	
<220> <221> CDS	
<222> 154675	
<221> sig_peptide	
<222> 154498	
<223> Von Heijne matrix score 4.8	
seq PLRLLNLLILIEG/GV	
<221> polyA signal	
<222> 819824	
<221> polyA_site	
<222> 838849	
<400> 140	
cccctatctc cagacctcat tcgcaatgaa gtagaatgtc tgaaagcaga tttcaaccac	60
agaatcaagg aggttetett caacteette tteagtgeet actatgttge attteteece etgtgttttg tgaagagtae ecagtaetat gae atg ege tgg tea tgt gag eac	
Cididificial idaddadiac coagractae gae acg ege egg eea egg eea	120 174
Met Arg Trp Ser Cys Glu His	174
Met Arg Trp Ser Cys Glu His -115 -110	174
Met Arg Trp Ser Cys Glu His -115 -110 ctc qtt atq qtq tqq atc aat gct ttt gtc atg ctc acc acg caa ctg	
Met Arg Trp Ser Cys Glu His -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu -105 -100 -95	174 222
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg	174
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg  Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg  Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85 -80	174 222 270
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg  Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg  Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85 -80  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag	174 222
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg  Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg  Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85 -80	174 222 270
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg  Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg  Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85 -80  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag  Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75 -70 -65  cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg	174 222 270
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85 -80  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75 -70 -65  cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg His Ile Trp Ser Glu Asn Thr Ile Trp Pro Gln Gly Val Leu Val Arg	174 222 270 318
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg  Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg  Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85 -80  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag  Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75 -70 -65  cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg  His Ile Trp Ser Glu Asn Thr Ile Trp Pro Gln Gly Val Leu Val Arg  -60 -55 -50 -45  cac agc agg gtg	174 222 270 318
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75 -70 -65  cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg His Ile Trp Ser Glu Asn Thr Ile Trp Pro Gln Gly Val Leu Val Arg  -60 -55 -50 -45  cac agc aga tgt tta tat aga gcc atg ggg cct tac aac gtg gca gtg His Ser Arg Cys Leu Tyr Arg Ala Met Gly Pro Tyr Asn Val Ala Val	174 222 270 318 366
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75  cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg His Ile Trp Ser Glu Asn Thr Ile Trp Pro Gln Gly Val Leu Val Arg  -60 -55 -50 -45  cac agc aga tgt tta tat aga gcc atg ggg cct tac aac gtg gca gtg His Ser Arg Cys Leu Tyr Arg Ala Met Gly Pro Tyr Asn Val Ala Val  -40 -35 -30  cct tca gat gta tct cat gcc cgc ttt tat ttt tta ttt cat cga cca	174 222 270 318 366
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85 -80  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75 cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg His Ile Trp Ser Glu Asn Thr Ile Trp Pro Gln Gly Val Leu Val Arg  -60 -55 -50 -45 cac agc aga tgt tta tat aga gcc atg ggg cct tac aac gtg gca gtg His Ser Arg Cys Leu Tyr Arg Ala Met Gly Pro Tyr Asn Val Ala Val  -40 -35 -30 cct tca gat gta tct cat gcc cgc ttt tat ttc tta ttt cat cga cca Pro Ser Asp Val Ser His Ala Arg Phe Tyr Phe Leu Phe His Arg Pro	174 222 270 318 366 414
Met Arg Trp Ser Cys Glu His  -115 -110  ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105 -100 -95  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90 -85  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75  cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg His Ile Trp Ser Glu Asn Thr Ile Trp Pro Gln Gly Val Leu Val Arg  -60 -55 -50 -45  cac agc aga tgt tta tat aga gcc atg ggg cct tac aac gtg gca gtg His Ser Arg Cys Leu Tyr Arg Ala Met Gly Pro Tyr Asn Val Ala Val  -40 -35 -30  cct tca gat gta tct cat gcc cgc ttt tat ttc tta ttt cat cga cca Pro Ser Asp Val Ser His Ala Arg Phe Tyr Phe Leu Phe His Arg Pro  -20 -15	174 222 270 318 366 414
## Arg Trp Ser Cys Glu His    Ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg   Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu   -105	174 222 270 318 366 414 462
ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu  -105  ttg cca tcc aaa tac tgt gat ttg cta cat aaa tca gct gct cac ctg Leu Pro Ser Lys Tyr Cys Asp Leu Leu His Lys Ser Ala Ala His Leu  -90  ggc aag tgg cag aag ttg gaa cat ggg tcc tac agc aat gct cca cag Gly Lys Trp Gln Lys Leu Glu His Gly Ser Tyr Ser Asn Ala Pro Gln  -75  cac att tgg tca gaa aat aca ata tgg cct caa ggg gtg ctg gtg cgg His Ile Trp Ser Glu Asn Thr Ile Trp Pro Gln Gly Val Leu Val Arg  -60  -55  cac agc aga tgt tta tat aga gcc atg ggg cct tac aac gtg gca gtg His Ser Arg Cys Leu Tyr Arg Ala Met Gly Pro Tyr Asn Val Ala Val  -40  cct tca gat gta tct cat gcc cgc ttt tat ttc tta ttt cat cga cca Pro Ser Asp Val Ser His Ala Arg Phe Tyr Phe Leu Phe His Arg Pro  -25  tta agg ctg tta aat ctg ctc atc ctt att gag ggc ggt gtc gtc ttc Leu Arg Leu Leu Asn Leu Leu Ile Leu Ile Glu Gly Gly Val Val Phe  -10  -10  -10  -10  -10  -10  -10  -1	174 222 270 318 366 414 462
## Arg Trp Ser Cys Glu His    Ctc gtt atg gtg tgg atc aat gct ttt gtc atg ctc acc acg caa ctg   Leu Val Met Val Trp Ile Asn Ala Phe Val Met Leu Thr Thr Gln Leu   -105	174 222 270 318 366 414 462 510

5 10 15 20 tcc atg gct ctc atc ctc ttc tgc aac tac tat gtt tta ttt aaa ctt Ser Met Ala Leu Ile Leu Phe Cys Asn Tyr Tyr Val Leu Phe Lys Leu 25 25 30 30 35 ctc cgg gac aga ata gta tta ggc agg gca tac tcc tac cca ctc aac Leu Arg Asp Arg Ile Val Leu Gly Arg Ala Tyr Ser Tyr Pro Leu Asn 40 45 50	606 654
agt tat gaa ctc aag gca aac taagctgcct ctcaacaatg agggagaact Ser Tyr Glu Leu Lys Ala Asn 55 cagataaaaa tattttcata cgttctattt ttttcttgtg atttttataa atatttaaga tgttttatat tttgtatact attatgtttt gaaagtcggg aagagtaagg gatattaaat	765 825
gtatccgtaa acaaaaaaa aaaa	849

<211> 155 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -31..-1 <400> 141 Met Phe Thr Ser Thr Gly Ser Ser Gly Leu Tyr Lys Ala Pro Leu Ser -25 Lys Ser Leu Leu Leu Val Pro Ser Ala Leu Ser Leu Leu Leu Ala Leu - 5 -10 Leu Leu Pro His Cys Gln Lys Pro Phe Val Tyr Asp Leu His Ala Val Lys Asn Asp Phe Gln Ile Trp Arg Leu Ile Cys Gly Arg Ile Ile Cys 25 Leu Asp Leu Lys Asp Thr Phe Cys Ser Ser Leu Leu Ile Tyr Asn Phe 40 Arg Ile Phe Glu Arg Arg Tyr Gly Ser Arg Lys Phe Ala Ser Phe Leu 60 -55 Leu Gly Thr Trp Val Leu Ser Ala Leu Phe Asp Phe Leu Leu Ile Glu 70 Ala Met Gln Tyr Phe Phe Gly Ile Thr Ala Ala Ser Asn Leu Pro Ser 90 85 Gly Leu Ile Phe Cys Cys Ala Phe Cys Ser Glu Thr Lys Leu Phe Leu 105 Ser Arg Gln Ala Met Ala Glu Asn Phe Ser Ile

120

<210> 142 <211> 55 <212> PRT <213> Homo sapiens

<210> 141

55

```
<210> 143
<211> 67
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -20..-1
<400> 143
Met Ser Arg Asn Leu Arg Thr Ala Leu Ile Phe Gly Gly Phe Ile Ser
                -15
                                        -10
Leu Ile Gly Ala Ala Phe Tyr Pro Ile Tyr Phe Arg Pro Leu Met Arg
Leu Glu Glu Tyr Lys Lys Glu Gln Ala Ile Asn Arg Ala Gly Ile Val
                           20
Gln Glu Asp Val Gln Pro Pro Gly Leu Lys Val Trp Ser Asp Pro Phe
Gly Arg Lys
45
<210> 144
<211> 198
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 144
Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala Leu Ala Met Val Thr
                                           -10
                        -15
Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro Glu Leu Ala Gln His
Glu Glu Leu Thr Leu Leu Phe His Gly Thr Leu Gln Leu Gly Gln Ala
                                20
Leu Asn Gly Val Tyr Arg Thr Thr Glu Gly Trp Leu Thr Lys Ala Arg
                            35
Asn Ser Leu Gly Leu Tyr Gly Arg Thr Ile Glu Leu Leu Gly Gln Glu
                                            55
                        50
Val Ser Arg Gly Arg Asp Ala Ala Gln Glu Leu Arg Ala Ser Leu Leu
                                        70
                    65
Glu Thr Gln Met Glu Glu Asp Ile Leu Gln Leu Gln Ala Glu Ala Thr
                                    85
                80
Ala Glu Val Leu Gly Glu Val Ala Gln Ala Gln Lys Val Leu Arg Asp
                                100
            95
Ser Val Gln Arg Leu Glu Val Gln Leu Arg Ser Ala Trp Leu Gly Pro
                                                120
        110
                            115
Ala Tyr Arg Glu Phe Glu Val Leu Lys Ala His Ala Asp Lys Gln Ser
                                            135
                        130
His Ile Leu Trp Ala Leu Thr Gly His Val Gln Arg Gln Arg Arg Glu
                    145
                                       150
```

Met Val Ala Gln Gln His Arg Leu Arg Gln Ile Gln Glu Arg Leu His

160

Thr Ala Ala Leu Pro Ala

```
<210> 145
<211> 135
<212> PRT
<213> Homo sapiens
<220>
               . . ---
<221> SIGNAL
<222> -25..-1
<400> 145
Met Ser Leu Arg Asn Leu Trp Arg Asp Tyr Lys Val Leu Val Val Met
                         -15
            . -20
Val Pro Leu Val Gly Leu Ile His Leu Gly Trp Tyr Arg Ile Lys Ser
Ser Pro Val Phe Gln Ile Pro Lys Asn Asp Asp Ile Pro Glu Gln Asp
                          15
Ser Leu Gly Leu Ser Asn Leu Gln Lys Ser Gln Ile Gln Gly Lys Xaa
                      30
Ala Gly Leu Gln Ser Ser Gly Lys Glu Ala Ala Leu Asn Leu Ser Phe
                  45
Ile Ser Lys Glu Glu Met Lys Asn Thr Ser Trp Ile Arg Lys Asn Trp
              60
Leu Leu Val Ala Gly Ile Ser Phe Ile Gly Asp His Leu Gly Thr Tyr
                               80-
Phe Leu Gln Arg Ser Ala Lys Gln Ser Val Lys Phe Gln Ser Gln Ser
                          95
Lys Gln Lys Ser Ile Glu Glu
<210> 146
<211> 255
<212> PRT
<213> Homo sapiens
```

<211> 255 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -70..-1

<400> 146 Met Gln Gln Lys Glu Gln Gln Phe Arg Glu Trp Phe Leu Lys Glu Phe -60 -65 -70 Pro Gln Ile Arg Trp Lys Ile Gln Glu Ser Ile Glu Arg Leu Arg Val -45 ~50 Ile Ala Asn Glu Ile Glu Lys Val His Arg Gly Cys Val Ile Ala Asn -25 -30 -35 Val Val Ser Gly Ser Thr Gly Ile Leu Ser Val Ile Gly Val Met Leu -15 -10 -20 Ala Pro Phe Thr Ala Gly Leu Ser Leu Ser Ile Thr Ala Ala Gly Val 5 Gly Leu Gly Ile Ala Ser Ala Thr Ala Gly Ile Ala Ser Ser Ile Val 20 15 Glu Asn Thr Tyr Thr Arg Ser Ala Glu Leu Thr Ala Ser Arg Leu Thr 35 Ala Thr Ser Thr Asp Gln Leu Glu Ala Leu Arg Asp Ile Leu His Asp 50 . Ile Thr Pro Asn Val Leu Ser Phe Ala Leu Asp Phe Asp Glu Ala Thr Lys Met Ile Ala Asn Asp Val His Thr Leu Arg Arg Ser Lys Ala Thr 90
Val Gly Arg Pro Leu Ile Ala Trp Arg Tyr Val Pro Ile Asn Val Val Val 105
Glu Thr Leu Arg Thr Arg Gly Ala Pro Thr Arg Ile Val Arg Lys Val 115
Ala Arg Asn Leu Gly Lys Ala Thr Ser Gly Val Leu Val Val Val Leu Asp 125
Val Val Asn Leu Val Gln Asp Ser Leu Asp Leu His Lys Gly Glu Lys 145
Ser Glu Ser Ala Glu Leu Thr His Ile His Gln Ser Leu Lys Ala Gly Lys Ala Gly Lys Lys Leu Lys Ala Gly Lys Leu Val Clu Gly Lys Lys Leu Asp 165
Asn Leu Asn Glu Leu Thr His Ile His Gln Ser Leu Lys Ala Gly Lys 185

<210> 147
<211> 59
<212> PRT
<213> Homo sapiens

<220>
<221> SIGNAL
<222> -49..-1

<400> 147
Met Pro Gly Thr Glu Val Let

Met Pro Gly Thr Glu Val Leu Glu Gly Ala Thr Asp Gly Leu Ala Ala
-45

Ile Asn Leu Leu Lys Trp Ile Lys Thr Leu Gly Gly Ser Val Ile Ser
-30

Met Ile Val Leu Leu Ile Cys Val Val Cys Leu Tyr Ile Val Cys Arg
-15

Cys Gly Ser His Leu Trp Arg Glu Ser His His
1

1

<210> 148 <211> 180 <212> PRT <213> Homo sapiens

<400> 148 Met Cys Ile Ser Gly Leu Cys Gln Ile Val Gly Cys Asp His Gln Leu Gly Ser Thr Val Lys Glu Asp Asn Cys Gly Val Cys Asn Gly Asp Gly Ser Thr Cys Arg Leu Val Arg Gly Gln Tyr Lys Ser Gln Leu Ser Ala Thr Lys Ser Asp Asp Thr Val Val Ala Ile Pro Tyr Gly Ser Arg His 55 Ile Arg Leu Val Leu Lys Gly Pro Asp His Leu Tyr Leu Glu Thr Lys 75 70 Thr Leu Gln Gly Thr Lys Gly Glu Asn Ser Leu Ser Ser Thr Gly Thr 90 Phe Leu Val Asp Asn Ser Ser Val Asp Phe Gln Lys Phe Pro Asp Lys 105 Glu Ile Leu Arg Met Ala Gly Pro Leu Thr Ala Asp Phe Ile Val Lys 120 Ile Arg Asn Ser Gly Ser Ala Asp Ser Thr Val Gln Phe Ile Phe Tyr

<210> 149 <211> 162 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -23..-1 <400> 149 Met Gly Asp Lys Ile Trp Leu Pro Phe Pro Val Leu Leu Ala Ala -20 -15 Leu Pro Pro Val Leu Leu Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu Asp Ser Asp Phe Thr Phe Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe Tyr Gln Pro Met Pro Leu Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val Leu Asp Gly Ala Gly Leu Asp Ile Asp Phe His Leu Ala Ser Pro Glu 50 Gly Lys Thr Leu Val Phe Glu Gln Arg Lys Ser Asp Gly Val His Thr 65 Val Glu Thr Glu Val Gly Asp Tyr Met Phe Cys Phe Asp Asn Thr Phe

Val Glu Thr Glu Val Gly Asp Tyr Met Phe Cys Phe Asp Ash Thr Phe
75 80 85

Ser Thr Ile Ser Glu Lys Val Ile Phe Phe Glu Leu Ile Pro Asp Ash
90 95 100 105

Met Gly Glu Gln Ala Gln Glu Gln Glu Asp Trp Lys Lys Tyr Ile Thr

Gly Thr Asp Ile Leu Asp Met Lys Leu Glu Asp Ile Leu Val Ser Met

120
130
135

Val Phe

<210> 150 <211> 120 <212> PRT <213> Homo sapiens

<220>
<221> SIGNAL
<222> -23..-1

Leu Asp Gly Ala Gly Leu Asp Ile Asp Phe His Leu Ala Ser Pro Glu
45
Gly Lys Thr Leu Val Phe Glu Gln Arg Lys Ser Asp Gly Val His Thr
60
Cys Ile Arg Ser Lys Asn Gly Pro Gly Thr Ala Val His Ala Tyr Asn
75
Pro Ser Thr Phe Arg Gly Gln Val
90

<210> 151 <211> 7 <212> PRT <213> Homo sapiens <400> 151 Met Val Glu Met Thr Gly Val 1 5

<210> 152 <211> 199 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -42..-1

<400> 152 Met Asp Gly Gln Lys Lys Asn Trp Lys Asp Lys Val Val Asp Leu Leu -35 Tyr Trp Arg Asp Ile Lys Lys Thr Gly Val Val Phe Gly Ala Ser Leu -15 -20 Phe Leu Leu Ser Leu Thr Val Phe Ser Ile Val Ser Val Thr Ala - 5 Tyr Ile Ala Leu Ala Leu Leu Ser Val Thr Ile Ser Phe Arg Ile Tyr 15 10 Lys Gly Val Ile Gln Ala Ile Gln Lys Ser Asp Glu Gly His Pro Phe 30 Arg Ala Tyr Leu Glu Ser Glu Val Ala Ile Ser Glu Glu Leu Val Gln 45 Lys Tyr Ser Asn Ser Ala Leu Gly His Val Asn Cys Thr Ile Lys Glu Leu Arg Arg Leu Phe Leu Val Asp Asp Leu Val Asp Ser Leu Lys Phe 80 Ala Val Leu Met Trp Val Phe Thr Tyr Val Gly Ala Leu Phe Asn Gly 95 Leu Thr Leu Leu Ile Leu Ala Leu Ile Ser Leu Phe Ser Val Pro Val 110 105 Ile Tyr Glu Arg His Gln Ala Gln Ile Asp His Tyr Leu Val Leu Ala 130 125 120 Asn Lys Asn Val Lys Asp Ala Met Ala Lys Ile Gln Ala Lys Ile Pro 140 Gly Leu Lys Arg Lys Ala Glu 155

<211> 43 <212> PRT <213> Homo sapiens

<210> 154 <211> 50 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1

<210> 155 <211> 153 <212> PRT <213> Homo sapiens

<400> 155 Thr Val Pro Leu Leu Glu Pro Ala Asp His Ala Arg Gly Arg Ala 10 His Val His Leu Pro Glu Asn Val Arg Ser Gln Ser Pro Gly His Val 25 Arg Arg Gly Arg Ser Gly Ala Gln Val Leu Pro Thr Gly Pro Asp Glu 45 40 Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 60 55 Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 70 Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 105 Trp Gly Glu Leu Leu Arg Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly 120 Glu Leu Lys Thr Glu Leu Leu Gly Leu Lys Glu Arg Lys His Lys Pro 135 Gln Val Ser Gln Gln Glu Glu Leu Lys

<210> 157 <211> 87 <212> PRT <213> Homo sapiens

<210> 156

<210> 158
<211> 250
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -85..-1

 4400> 158

 Met Ser Ala Glu Val Lys Val Thr Gly Gln Asn Gln Glu Gln Phe Leu -85
 -80
 -75
 -75
 -75
 -70

 Leu Leu Ala Lys Ser Ala Lys Gly Ala Lys Gly Ala Leu Leu Ala Thr Leu Ile His -65
 -65
 -60
 -60
 -55
 -55

 Gln Val Leu Glu Ala Pro Gly Val Tyr Val Phe Gly Glu Leu Leu Asp -50
 -45
 -45
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40
 -40

Leu Arg Asn Val Arg Gln Leu Glu Asp Leu Val Ile Glu Ala Val Tyr 20 15 Ala Asp Val Leu Arg Gly Ser Leu Asp Gln Arg Asn Gln Arg Leu Glu Val Asp Tyr Ser Ile Gly Arg Asp Ile Gln Arg Gln Asp Leu Ser Ala Ile Ala Arg Thr Leu Gln Glu Trp Cys Val Gly Cys Glu Val Val Leu Ser Gly Ile Glu Glu Gln Val Ser Arg Ala Asn Gln His Lys Glu Gln 85 Gln Leu Gly Leu Lys Gln Gln Ile Glu Ser Glu Val Ala Asn Leu Lys 100 Lys Thr Ile Lys Val Thr Thr Ala Ala Ala Ala Ala Thr Ser Gln 115 Asp Pro Glu Gln His Leu Thr Glu Leu Arg Glu Pro Ala Pro Gly Thr 135 130 Asn Gln Arg Gln Pro Ser Lys Lys Ala Ser Lys Gly Lys Gly Leu Arg 145 Gly Ser Ala Lys Ile Trp Ser Lys Ser Asn 160

<210> 159 <211> 24 <212> PRT <213> Homo sapiens

<210> 160 <211> 228 <212> PRT <213> Homo sapiens

<400> 160 Met Pro Thr Asn Cys Ala Ala Ala Gly Cys Ala Thr Thr Tyr Asn Lys 10 His Ile Asn Ile Ser Phe His Arg Phe Pro Leu Asp Pro Lys Arg Arg Lys Glu Trp Val Arg Leu Val Arg Arg Lys Asn Phe Val Pro Gly Lys 40 His Thr Phe Leu Cys Ser Lys His Phe Glu Ala Ser Cys Phe Asp Leu Thr Gly Gln Thr Arg Arg Leu Lys Met Asp Ala Val Pro Thr Ile Phe Asp Phe Cys Thr His Ile Lys Ser Met Lys Leu Lys Ser Arg Asn Leu 90 Leu Lys Lys Asn Asn Ser Cys Ser Pro Ala Gly Pro Ser Ser Leu Lys 105 Ser Asn Ile Ser Ser Gln Gln Val Leu Leu Glu His Ser Tyr Ala Phe 120 Arg Asn Pro Met Glu Ala Lys Lys Arg Ile Ile Lys Leu Glu Lys Glu 135 Ile Ala Ser Leu Arg Arg Lys Met Lys Thr Cys Leu Gln Lys Glu Arg

<210> 161 <211> 86 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -20..-1 <400> 161 Met Asn Leu His Phe Pro Gln Trp Phe Val His Ser Ser Ala Leu Gly -15 -10 Leu Val Leu Ala Pro Pro Phe Ser Ser Pro Gly Thr Asp Pro Thr Phe 5 1 Pro Cys Ile Tyr Cys Arg Leu Leu Asn Met Ile Met Thr Arg Leu Ala 20 Phe Ser Phe Ile Thr Cys Leu Cys Pro Asn Leu Lys Glu Val Cys Leu 40 35 Ile Leu Pro Glu Lys Asn Cys Asn Ser Arg His Ala Gly Phe Val Gly

<210> 162 <211> 44 <212> PRT <213> Homo sapiens

Pro Ala Lys Leu Arg Gln

50

<210> 163
<211> 314
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -58..-1

<400> 163 Met Gln Asn Val Ile Asn Thr Val Lys Gly Lys Ala Leu Glu Val Ala -55 -50 Glu Tyr Leu Thr Pro Val Leu Lys Glu Ser Lys Phe Arg Glu Thr Gly -35 Val Ile Thr Pro Glu Glu Phe Val Ala Ala Gly Asp His Leu Val His -15 -20 His Cys Pro Thr Trp Gln Trp Ala Thr Gly Glu Glu Leu Lys Val Lys -5 Ala Tyr Leu Pro Thr Gly Lys Gln Phe Leu Val Thr Lys Asn Val Pro 15 Cys Tyr Lys Arg Cys Lys Gln Met Glu Tyr Ser Asp Glu Leu Glu Ala 30 Ile Ile Glu Glu Asp Asp Gly Asp Gly Gly Trp Val Asp Thr Tyr His 45 Asn Thr Gly Ile Thr Gly Ile Thr Glu Ala Val Lys Glu Ile Thr Leu 60 Glu Asn Lys Asp Asn Ile Arg Leu Gln Asp Cys Ser Ala Leu Cys Glu 80 75 Glu Glu Glu Asp Glu Asp Glu Gly Glu Ala Ala Asp Met Glu Glu Tyr 95 Glu Glu Ser Gly Leu Leu Glu Thr Asp Glu Ala Thr Leu Asp Thr Arg 115 110 Lys Ile Val Glu Ala Cys Lys Ala Lys Thr Asp Ala Gly Gly Glu Asp 130 125 Ala Ile Leu Gln Thr Arg Thr Tyr Asp Leu Tyr Ile Thr Tyr Asp Lys 145 140 Tyr Tyr Gln Thr Pro Arg Leu Trp Leu Phe Gly Tyr Asp Glu Gln Arg 160 155 Gln Pro Leu Thr Val Glu His Met Tyr Glu Asp Ile Ser Gln Asp His 175 170 Val Lys Lys Thr Val Thr Ile Glu Asn His Pro His Leu Pro Pro Pro 190 Pro Met Cys Ser Val His Pro Cys Arg His Ala Glu Val Met Lys Lys 210 205 Ile Ile Glu Thr Val Ala Glu Gly Gly Glu Leu Gly Val His Met 225 220 Tyr Leu Leu Ile Phe Leu Lys Phe Val Gln Ala Val Ile Pro Thr Ile 240 Glu Tyr Asp Tyr Thr Arg His Phe Thr Met

<210> 164 <211> 89 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -80..-1

WO 99/31236

-30 -25 -20

Gln Leu Gly Arg Gly Leu Leu Ser Ala Cys Ala Pro Trp Gly Asp Gly
-15 -10 -5

Ser Thr Gln Pro Val Pro Leu Cys Ser
1 5

<210> 165 <211> 98 <212> PRT <213> Homo sapiens

<220>
<221> SIGNAL
<222> -15..-1

<210> 166 <211> 92 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -36..-1

<210> 167
<211> 351
<212> PRT

<213> Homo sapiens <220>

<221> SIGNAL <222> -16..-1

<400> 167 Met Val Pro Phe Ile Tyr Leu Gln Ala His Phe Thr Leu Cys Ser Gly -10 Trp Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val Tyr Val Pro Tyr 10 Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp Leu Val Ser Ile 25 Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile Ala Ala Ile Thr 40 Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp Glu Gly Ile Leu 55 Gly Leu Ala Tyr Ala Glu Ile Ala Arg Pro Asp Asp Ser Pro Glu Pro 70 Phe Phe Asp Ser Leu Val Lys Gln Thr His Val Pro Asn Leu Phe Ser 90 85 Leu Gln Leu Cys Gly Ala Gly Phe Pro Leu Asn Gln Ser Glu Val Leu 105 100 Ala Ser Val Gly Gly Ser Met Ile Ile Gly Gly Ile Asp His Ser Leu 125 120 115 Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr 135 140 Glu Val Ile Ile Val Arg Val Glu Ile Asn Gly Gln Asp Leu Lys Met 155 150 Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr 170 165 Thr Asn Leu Arg Leu Pro Lys Lys Val Phe Glu Ala Ala Val Lys Ser 185 Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu 200 205 Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile 220 215 Phe Pro Val Ile Ser Leu Tyr Leu Met Gly Glu Val Thr Asn Gln Ser 235 230 Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp 245 250 Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser 265 . 260 Ser Thr Gly Thr Val Met Gly Ala Val Ile Met Glu Gly Phe Tyr Val 280 285 275 Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala Val Ser Ala Cys 300 295 His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu Gly Pro Phe Cys 315 310 His Leu Gly His Gly Arg Leu Trp Leu Gln His Ser Thr Asp Arg

330

<210> 168 <211> 138 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -47..-1

<400> 168 Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly Ile Asp Leu -40 -35 -45 Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro Phe Val Ser -25 -30 Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys Ala Cys Ile -10 Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu His Leu Leu 10 Ala Gly Leu Cys Thr Leu Gly Ser Val Ser Cys Tyr Val Ala Gly Ile 25 Glu Leu Leu His Gln Lys Leu Glu Leu Pro Asp Asn Val Ser Gly Glu 4.5 Phe Gly Trp Ser Phe Cys Leu Ala Cys Val Ser Ala Pro Leu Gln Phe 60 Met Ala Ser Ala Leu Phe Ile Trp Ala Ala His Thr Asn Arg Arg Glu 75 Tyr Thr Leu Met Lys Ala Tyr Arg Val Ala

<210> 169
<211> 101
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -73..-1

<400> 169 Met Asn Leu Glu Arg Val Ser Asn Glu Glu Lys Leu Asn Leu Cys Arg -65 -70 Lys Tyr Tyr Leu Gly Gly Phe Ala Phe Leu Pro Phe Leu Trp Leu Val -50 -55 Asn Ile Phe Trp Phe Tyr Arg Glu Ala Phe Leu Val Pro Ala Tyr Thr -35 -30 Glu Gln Ser Gln Ile Lys Gly Tyr Val Trp Arg Ser Ala Val Gly Phe -15 -20 Leu Phe Trp Val Ile Val Leu Thr Ser Trp Ile Thr Ile Phe Gln Ile 1 ~5 Tyr Arg Pro Arg Trp Gly Ala Leu Gly Asp Tyr Leu Ser Phe Thr Ile 15 10 Pro Leu Gly Thr Pro 25

<210> 170 <211> 252 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -68..-1

Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -45 Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro -10 -15 Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His 20 Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu 35 Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe Arg Glu Asn Val Leu Arg Asn Leu Ala Asp Lys Ala Phe Asp Arg Pro 8.5 Ile Cys Glu Ala Leu Leu Asp Gln Arg Phe Phe Asn Gly Ile Gly Asn 100 Tyr Leu Arg Ala Glu Ile Leu Tyr Arg Leu Lys Ile Pro Pro Phe Glu 115 120 Lys Ala Arg Ser Val Leu Glu Ala Leu Gln Gln His Arg Pro Ser Pro 135 130 Glu Leu Thr Leu Ser Gln Lys Ile Arg Thr Lys Leu Gln Asn Ser Asp 150 145 Leu Leu Glu Leu Cys His Ser Val Pro Lys Glu Val Val Gln Leu Gly 165 Glu Ala Lys Asp Gly Ser Asn Leu Cys Phe Ser Lys

<210> 171 <211> 350 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -68..-1

<400> 171 Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu -60 Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -45 -50 Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser -25 -30 Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro -10 -20 -15 Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His 20 15 Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu 3.5 Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys 50 Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe 70 Arg Leu Lys Ile Pro Pro Phe Glu Lys Ala Arg Ser Val Leu Glu Ala

```
85
          80
Leu Gln Gln His Arg Pro Ser Pro Glu Leu Thr Leu Ser Gln Lys Ile
                                          105
                         100
Arg Thr Lys Leu Gln Asn Pro Asp Leu Leu Glu Leu Cys His Ser Val
                                       120
                     115
Pro Lys Glu Val Asp Gln Leu Gly Gly Arg Gly Tyr Gly Ser Glu Ser
                                   135
                 130
Gly Glu Glu Asp Phe Ala Ala Phe Arg Ala Trp Leu Arg Cys Tyr Gly
                               150
              145
Met Pro Gly Met Ser Ser Leu Gln Asp Arg His Gly Arg Thr Ile Trp
                                               170
                            165
          160
Phe Gln Gly Asp Pro Gly Pro Leu Ala Pro Lys Gly Arg Lys Ser Arg
                        180
                                           185
      175
Lys Lys Lys Ser Lys Ala Thr Gln Leu Ser Pro Glu Asp Arg Val Glu
                             200
                     195
Asp Ala Leu Pro Pro Ser Lys Ala Pro Ser Lys Thr Arg Arg Ala Lys
                         215
                 210
Arg Asp Leu Pro Lys Arg Thr Ala Thr Gln Arg Pro Glu Gly Thr Ser
                      230
             225
Leu Gln Gln Asp Pro Glu Ala Pro Thr Val Pro Lys Lys Gly Arg Arg
                   245
Lys Gly Arg Gln Ala Ala Ser Gly His Cys Arg Pro Arg Lys Val Lys
              260
Ala Asp Ile Pro Ser Leu Glu Pro Glu Gly Thr Ser Ala Ser
             275
```

<210> 172 <211> 390 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -68..-1 <400> 172 Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu -60 -65 Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -45 Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser -25 -30 Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro -15 -10 Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His 20 15 Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu 35 Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys 50 Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe 65 Arg Glu Asn Val Leu Arg Asn Leu Ala Asp Lys Ala Phe Asp Arg Pro 90 Ile Cys Glu Ala Leu Leu Asp Gln Arg Phe Phe Asn Gly Ile Gly Asn 105 100 Tyr Leu Arg Ala Glu Ile Leu Tyr Arg Leu Lys Ile Pro Pro Phe Glu

```
Lys Ala Arg Ser Val Leu Glu Ala Leu Gln Gln His Arg Pro Ser Pro
                            135 140
              130
Glu Leu Thr Leu Ser Gln Lys Ile Arg Thr Lys Leu Gln Asn Pro Asp
                               150
              145
Leu Leu Glu Leu Cys His Ser Val Pro Lys Glu Val Val Gln Leu Gly
                           165
          160
Gly Arg Gly Tyr Gly Ser Glu Ser Gly Glu Glu Asp Phe Ala Ala Phe
                        180
       175
Arg Ala Trp Leu Arg Cys Tyr Gly Met Pro Gly Met Ser Ser Leu Gln
                    195
Asp Arg His Gly Arg Thr Ile Trp Phe Gln Gly Asp Pro Gly Pro Leu
                210
                                  215
Ala Pro Lys Gly Arg Lys Ser Arg Lys Lys Lys Ser Lys Ala Thr Gln
                             230
              225
Leu Ser Pro Glu Asp Arg Val Glu Asp Ala Leu Pro Pro Ser Lys Ala
                           245
Pro Ser Arg Thr Arg Arg Ala Lys Arg Asp Leu Pro Lys Arg Thr Ala
                              265
                     260
Thr Gln Arg Pro Glu Gly Thr Ser Leu Gln Gln Asp Pro Glu Ala Pro
                             280
                    275
Thr Val Pro Lys Lys Gly Arg Arg Lys Gly Arg Gln Ala Ala Ser Gly
                            295
                 290
His Cys Arg Pro Arg Lys Val Lys Ala Asp Ile Pro Ser Leu Glu Pro
                     310
           305
Glu Gly Thr Ser Ala Ser
          320
```

<210> 173 <211> 190 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -82..-1 <400> 173 Met Tyr Val Trp Pro Cys Ala Val Val Leu Ala Gln Tyr Leu Trp Phe -70 -75 His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly -55 -60 Val Ser Leu Pro Gly Ile Leu Thr Ala Lys Cys Gly Ala Glu Val Ile -40 -45 Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln -25 -30 Ser Cys Gln Met Asn Asn Leu Pro His Leu Gln Val Val Gly Leu Thr -10 -15 Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Asp Phe Glu Asp Ile 20 25 Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln Leu 40 35 Trp Ser Thr Tyr Gln Val Arg Ser Ala Asp Trp Ser Leu Glu Ala Leu 55 Leu Tyr Lys Trp Asp Met Lys Cys Val His Ile Pro Leu Glu Ser Phe 75 70 Asp Ala Asp Lys Glu Asp Ile Ala Glu Ser Thr Leu Pro Gly Arg His 85 Thr Val Glu Met Leu Val Ile Ser Phe Ala Lys Asp Ser Leu

100

105

```
<210> 174
<211> 285
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -232..-1
<400> 174
Met Gly Cys Val Phe Gln Ser Thr Glu Asp Lys Arg Ile Phe Lys Ile
   -230 -225 -220
Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu
         -210
                          -205
-215
Tyr Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg
      -195 -190 -185
-200
Val His Leu Met Gly Asp Asn Leu Cys Asn Asp Gly Ser Leu Leu Leu
           -180 -175 -170
Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg
      -165 -160 -155
Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val
    -150 -145 -140
Leu Pro Glu Glu Pro Lys Glu Leu Met Val His Val Gly Gly Leu Ile
 -135 -130 -125
Gln Met Gly Cys Val Phe Gln Ser Thr Glu Val Lys His Val Thr Lys
-120 -115 -110 -105
Val Glu Trp Ile Phe Ser Gly Arg Arg Ala Lys Glu Glu Ile Val Phe
         -100 -95 -90
Arg Tyr Tyr His Lys Leu Arg Met Ser Ala Glu Tyr Ser Gln Ser Trp
      -85
                         -80
Gly His Phe Gln Asn Arg Val Asn Leu Val Gly Asp Ile Phe Arg Asn
                                     -60
                     -65
Asp Gly Ser Ile Met Leu Gln Gly Val Arg Glu Ser Asp Gly Gly Asn
                                  -45
               -50
Tyr Thr Cys Ser Ile His Leu Gly Asn Leu Val Phe Lys Lys Thr Ile
                         -30
               -35
Val Leu His Val Ser Pro Glu Glu Pro Arg Thr Leu Val Thr Pro Ala
            -20
                         -15 -10
Ala Leu Arg Pro Leu Val Leu Gly Gly Asn Gln Leu Val Ile Ile Val
                      1
         -5
Gly Ile Val Cys Ala Thr Ile Leu Leu Leu Pro Val Leu Ile Leu Ile
                  15
Val Lys Lys Thr Cys Gly Asn Lys Ser Ser Val Asn Ser Thr Val Leu
             30
25
Val Lys Asn Thr Lys Lys Thr Asn Pro Lys Lys Lys
            45
```

<210> 175

<211> 153

<212> PRT

<213> Homo sapiens

<400> 175

Met Gly Cys Val Phe Gln Ser Thr Val Asp Lys Cys Ile Phe Lys Ile

1 5 10 15

Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu

<210> 176 <211> 49 <212> PRT <213> Homo sapiens

<210> 177
<211> 99
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -24..-1

```
<210> 178
<211> 95
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -37..-1
<400> 178
Met Ala Ser Pro Ala Val Asn Arg Trp Lys Arg Pro Arg Leu Lys Pro
                          -30
      -35
Val Trp Pro Arg Arg Leu Glu Ser Trp Leu Leu Leu Asp Ala Leu Leu
                                         -10
                      -15
Arg Leu Gly Asp Thr Lys Lys Lys Arg Gln Pro Glu Ala Ala Thr Lys
Ser Cys Val Arg Ser Ser Cys Gly Gly Pro Ser Gly Asp Gly Pro Pro
                             20
          1.5
Pro Cys Leu Gln Gln Pro Asp Pro Arg Ala Leu Ser Gln Ala Phe Ser
                         35
Arg Ser Phe Pro Leu Phe Pro Ser Leu Ala Gly Lys Ser Met Ile
                                 5.5
                    50 .
<210> 179
<211> 121
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -23..-1
<400> 179
Met Met Leu Pro Gln Trp Leu Leu Leu Phe Leu Leu Phe Phe Phe
                               -15
                                      -10
          -20
Leu Phe Leu Leu Thr Arg Gly Ser Leu Ser Pro Thr Lys Tyr Asn Leu
Leu Glu Leu Lys Glu Ser Cys Ile Arg Asn Gln Asp Cys Glu Thr Gly
Cys Cys Gln Arg Ala Pro Asp Asn Cys Glu Ser His Cys Ala Glu Lys
                                   35
Gly Ser Glu Gly Ser Leu Cys Gln Thr Gln Val Phe Phe Gly Gln Tyr
           45
Arg Ala Cys Pro Cys Leu Arg Asn Leu Thr Cys Ile Tyr Ser Lys Asn
    60
                           65
Glu Lys Trp Leu Ser Ile Ala Tyr Gly Arg Cys Gln Lys Ile Gly Arg
                   80
Gln Lys Leu Ala Lys Lys Met Phe Phe
                   95
```

<210> 180 <211> 59

<212> PRT

<213> Homo sapiens

<400> 180

Met Ile Leu Cys Phe Leu Leu Pro His His Arg Leu Gln Glu Ala Arg

10 15
Gln Ile Gln Val Leu Lys Met Leu Pro Arg Glu Lys Leu Arg Arg Arg
20 25 30
Glu Glu Arg Lys Gln Ile Asn Gly Lys Lys Glu Arg Thr Lys Tyr Glu
35 40 45
Thr Pro Arg Lys Arg Glu Gly Lys Lys Lys Lys Lys
50 55

<210> 181 <211> 86 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 181 Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp Cys -10 - 5 Asp Pro Arg Arg Ile His Ser Gln Asp Asp Val Pro Arg Ser Ser Ala 10 Ala Asp Thr Gly Ser Ala Met Gln Arg Arg Glu Ala Trp Ala Gly Trp 25 Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg Leu 40 45 Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu Gly 60 55 Tyr Arg Ile Cys Asp Leu

<210> 182 <211> 165 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -58..-1

70

<400> 182 Met Thr Arg Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro Ile -50 Pro Val Pro Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser Pro -30 -35 Val Arg Pro Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu Leu -15 -20 Asp Ser Val Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala Val - 5 -10 Phe Ser Thr Thr Gly Pro Ala Leu Leu Leu Leu Val Ser Phe Leu 15 10 Thr Phe Asp Leu Leu His Arg Pro Ala Gly His Thr Leu Pro Gln Arg 30 25 Lys Leu Leu Thr Arg Gly Gln Ser Gln Gly Ala Gly Glu Gly Pro Gly 45 50 Gln Gln Glu Ala Leu Leu Gln Met Gly Thr Val Ser Gly Gln Leu 65 Ser Leu Gln Asp Ala Leu Leu Leu Leu Leu Met Gly Leu Gly Pro Leu Leu Arg Ala Cys Gly Met Pro Leu Thr Leu Leu Gly Leu Ala Phe Cys 90 95 100

Leu His Pro Trp Ala 105

<210> 183
<211> 80
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -35..-1

<210> 184
<211> 73
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1

<210> 185 <211> 98 <212> PRT <213> Homo sapiens

<400> 185
Met Leu Gly Ala Glu Thr Glu Glu Lys Leu Phe Asp Ala Pro Leu Ser
1 5 10 15

 Ile
 Ser
 Lys
 Arg
 Glu
 Gln
 Leu
 Glu
 Gln
 Fro
 Fro
 Gln
 Fro
 Fro
 Gln
 Fro
 Fro
 Gln
 Fro
 Fro
 Fro
 Fro
 Fro
 Fro
 Gln
 Fro
 F

<210> 186 <211> 112 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 186 Met Glu Ser Arg Val Leu Leu Arg Thr Phe Cys Leu Ile Phe Gly Leu -15 Gly Ala Val Trp Gly Leu Gly Val Asp Pro Ser Leu Gln Ile Asp Val - 5 1 Leu Thr Glu Leu Glu Leu Gly Glu Ser Thr Thr Gly Val Arg Gln Val 20 Pro Gly Leu His Asn Gly Thr Lys Ala Phe Leu Phe Gln Asp Thr Pro 35 Arg Ser Ile Lys Ala Ser Thr Ala Thr Ala Glu Gln Phe Phe Gln Lys 50 Leu Arg Asn Lys His Glu Phe Thr Ile Leu Val Thr Leu Lys Gln Thr 70 65 His Leu Asn Ser Gly Val Ile Leu Ser Ile His His Leu Asp His Arg

<210> 187 <211> 70 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -44..-1

80

```
<210> 188
<211> 92
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -13..-1
<400> 188
Met Leu Phe Ser Leu Ser Leu Leu Ser Asn Leu Asn Gln Ile Gly Ser
                                - 5
           -10
Ser His Leu Asp Arg Pro His Ile Pro Gly Gln Ser Ala Gln Leu Phe
                        10
Ile Tyr Gln Met Ser Ser Gln Gln Leu Gln Gln Gln Pro Ser Ala Asn
                                       30
                    25
Lys Lys Ala Gly Lys Ile His Asn Thr Pro Phe Ala Asn Gln Leu Asn
                                    45
                40
Pro Thr Gln His Leu Ala Lys Pro Phe Gln Gln Ile Leu Pro Gly Arg
                               60
Gln Ser Gly Ser Leu Thr Ser Pro Phe Leu Ala Cys
```

<210> 189

<211> 207 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -42..-1 <400> 189 Met His Ile Leu Gln Leu Leu Thr Thr Val Asp Asp Gly Ile Gln Ala -35 Ile Val His Cys Pro Asp Thr Gly Lys Asp Ile Trp Asn Leu Leu Phe -15 -20 Asp Leu Val Cys His Glu Phe Cys Gln Ser Asp Asp Pro Pro Ile Ile 1 ~5 Leu Gln Glu Gln Lys Thr Val Leu Ala Ser Val Phe Ser Val Leu Ser 15 Ala Ile Tyr Ala Ser Gln Thr Glu Gln Glu Tyr Leu Lys Ile Glu Lys 30 Val Asp Leu Pro Leu Ile Asp Ser Leu Ile Arg Val Leu Gln Asn Met . 45 Glu Gln Cys Gln Lys Lys Pro Glu Asn Ser Ala Glu Ser Asn Thr Glu 65 60 Glu Thr Lys Arg Thr Asp Leu Thr Gln Asp Asp Leu His Leu Lys Ile 80 75 Leu Lys Asp Ile Leu Cys Glu Phe Leu Ser Asn Ile Phe Gln Ala Leu Thr Lys Glu Thr Val Ala Gln Gly Val Lys Glu Gly Gln Leu Ser Lys 115 110 Gln Lys Cys Ser Ser Ala Phe Gln Asn Leu Leu Pro Phe Tyr Ser Pro 125 130 Val Val Glu Asp Phe Ile Lys Ile Leu Arg Glu Val Asp Lys Ala Leu 140 Ala Asp Asp Leu Glu Lys Asn Phe Pro Ser Leu Lys Val Gln Thr

165 160 155

<210> 190 <211> 201 <212> PRT <213> Homo sapiens <400> 190 Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala Leu Lys Glu Lys Phe 10 Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe Gln Glu Ile Pro Lys 20 Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln Leu Glu Lys Ile Glu 70

Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val Asn His Leu Lys Ala 75 Asn Val Lys Ser Ala Ala Asp Leu Ile Ser Leu Pro Thr Thr Val Glu 90 Gly Leu Gln Lys Ser Val Ala Ser Ile Gly Asn Thr Leu Asn Ser Val 110 105 100 His Leu Ala Val Glu Ala Leu Gln Lys Thr Val Asp Glu His Lys Lys 120 125 115 Thr Met Glu Leu Leu Gln Ser Asp Met Asn Gln His Phe Leu Lys Glu 135 140

Thr Pro Gly Ser Asn Gln Ile Ile Pro Ser Pro Ser Ala Thr Ser Glu 155 150 Leu Asp Asn Lys Thr His Ser Glu Asn Leu Lys Gln Met Gly Asp Arg 170 165 Ser Ala Thr Leu Lys Arg Gln Ser Leu Asp Gln Val Thr Asn Arg Thr 185

Asp Thr Val Lys Ile Gln Lys Lys

<210> 191 <211> 379 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -37..-1

<400> 191 Met Pro His Ser Ser Leu His Pro Ser Ile Pro Cys Pro Arg Gly His -25 -30 Gly Ala Gln Lys Ala Ala Leu Val Leu Leu Ser Ala Cys Leu Val Thr -10 -15 Leu Trp Gly Leu Gly Glu Pro Pro Glu His Thr Leu Arg Tyr Leu Val 1 Leu His Leu Ala Ser Leu Gln Leu Gly Leu Leu Leu Asn Gly Val Cys 20 Ser Leu Ala Glu Glu Leu Arg His Ile His Ser Arg Tyr Arg Gly Ser 35 Tyr Trp Arg Thr Val Arg Ala Cys Leu Gly Cys Pro Leu Arg Arg Gly Ala Leu Leu Leu Ser Ile Tyr Phe Tyr Tyr Ser Leu Pro Asn Ala

70 Val Gly Pro Pro Phe Thr Trp Met Leu Ala Leu Leu Gly Leu Ser Gln 85 Ala Leu Asn Ile Leu Leu Gly Leu Lys Gly Leu Ala Pro Ala Glu Ile 100 Ser Ala Val Cys Glu Lys Gly Asn Phe Asn Val Ala His Gly Leu Ala 120 115 Trp Ser Tyr Tyr Ile Gly Tyr Leu Arg Leu Ile Leu Pro Glu Leu Gln 135 130 Ala Arg Ile Arg Thr Tyr Asn Gln His Tyr Asn Asn Leu Leu Arg Gly 150 145 Ala Val Ser Gln Arg Leu Tyr Ile Leu Leu Pro Leu Asp Cys Gly Val 165 160 Pro Asp Asn Leu Ser Met Ala Asp Pro Asn Ile Arg Phe Leu Asp Lys 180 175 Leu Pro Gln Gln Thr Gly Asp Arg Ala Gly Ile Lys Asp Arg Val Tyr 200 195 Ser Asn Ser Ile Tyr Glu Leu Leu Glu Asn Gly Gln Arg Ala Gly Thr 215 210 Cys Val Leu Glu Tyr Ala Thr Pro Leu Gln Thr Leu Phe Ala Met Ser 230 225 Gln Tyr Ser Gln Ala Gly Phe Ser Arg Glu Asp Arg Leu Glu Gln Ala 245 240 Lys Leu Phe Cys Arg Thr Leu Glu Asp Ile Leu Ala Asp Ala Pro Glu 260 255 Ser Gln Asn Asn Cys Arg Leu Ile Ala Tyr Gln Glu Pro Ala Asp Asp 275 Ser Ser Phe Ser Leu Ser Gln Glu Val Leu Arg His Leu Arg Gln Glu 290 Glu Lys Glu Glu Val Thr Val Gly Ser Leu Lys Thr Ser Ala Val Pro 310 305 Ser Thr Ser Thr Met Ser Gln Glu Pro Glu Leu Leu Ser Gly Met 325 320 Gly Lys Pro Leu Pro Leu Arg Thr Asp Phe Ser 335

<210> 192 <211> 112 <212> PRT

<213> Homo sapiens

 Add of the state of the st

<211> 43 <212> PRT <213> Homo sapiens

<210> 194 <211> 51 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -16..-1

<210 > 195 <211 > 244 <212 > PRT <213 > Homo sapiens <220 > <221 > SIGNAL <222 > -18..-1

<400> 195 Met Ala Asn Pro Lys Leu Leu Gly Leu Glu Leu Ser Glu Ala Glu Ala -10 Ile Gly Ala Asp Ser Ala Arg Phe Glu Glu Leu Leu Gln Ala Ser Lys Glu Leu Gln Gln Ala Gln Thr Thr Arg Pro Glu Ser Thr Gln Ile 25 20 Gln Pro Gln Pro Gly Phe Cys Ile Lys Thr Asn Ser Ser Glu Gly Lys 40 35 Val Phe Ile Asn Ile Cys His Ser Pro Ser Ile Pro Pro Pro Ala Asp 55 Val Thr Glu Glu Glu Leu Leu Gln Met Leu Glu Glu Asp Gln Ala Gly Phe Arg Ile Pro Met Ser Leu Gly Glu Pro His Ala Glu Leu Asp Ala 85 Lys Gly Gln Gly Cys Thr Ala Tyr Asp Val Ala Val Asn Ser Asp Phe 105 100 Tyr Arg Arg Met Gln Asn Ser Asp Phe Leu Arg Glu Leu Val Ile Thr

120 115 Ile Ala Arg Glu Gly Leu Glu Asp Ile Tyr Asn Leu Gln Leu Asn Pro 140 135 Glu Trp Arg Met Met Lys Asn Arg Pro Phe Met Gly Ser Ile Ser Gln 155 150 Gln Asn Ile Arg Ser Glu Gln Arg Pro Arg Ile Gln Glu Leu Gly Asp 170 165 Leu Tyr Thr Pro Ala Pro Gly Arg Ala Glu Ser Gly Pro Glu Lys Pro 185 180 His Leu Asn Leu Trp Leu Glu Ala Pro Asp Leu Leu Ala Glu Val 195 200 Asp Leu Pro Lys Leu Asp Gly Ala Leu Gly Leu Ser Leu Glu Ile Gly 210 Arg Thr Ala Trp 225

<210> 196
<211> 353
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -34..-1
<400> 196
Met Glu Arg Gly Let
-31
Thr Gly Trp Ala Gly

<400> 196 Met Glu Arg Gly Leu Lys Ser Ala Asp Pro Arg Asp Gly Thr Gly Tyr ~25 -30 Thr Gly Trp Ala Gly Ile Ala Val Leu Tyr Leu His Leu Tyr Asp Val ~ 5 -10 -15 Phe Gly Asp Pro Ala Tyr Leu Gln Leu Ala His Gly Tyr Val Lys Gln Ser Leu Asn Cys Leu Thr Lys Arg Ser Ile Thr Phe Leu Cys Gly Asp 25 20 Ala Gly Pro Leu Ala Val Ala Ala Val Leu Tyr His Lys Met Asn Asn 40 Glu Lys Gln Ala Glu Asp Cys Ile Thr Arg Leu Ile His Leu Asn Lys 55 Ile Asp Pro His Ala Pro Asn Glu Met Leu Tyr Gly Arg Ile Gly Tyr 70 Ile Tyr Ala Leu Leu Phe Val Asn Lys Asn Phe Gly Val Glu Lys Thr 85 Pro Gln Ser His Ile Gln Gln Ile Cys Glu Thr Ile Leu Thr Ser Gly 105 100 Glu Asn Leu Ala Arg Lys Arg Asn Phe Thr Ala Lys Ser Pro Leu Met 120 115 Tyr Glu Trp Tyr Gln Glu Tyr Tyr Val Gly Ala Ala His Gly Leu Ala 135 130 Gly Ile Tyr Tyr Leu Met Gln Pro Ser Leu Gln Val Ser Gln Gly 150 Lys Leu His Ser Leu Val Lys Pro Ser Val Asp Tyr Val Cys Gln Leu 170 165 Lys Phe Pro Ser Gly Asn Tyr Pro Pro Cys Ile Gly Asp Asn Arg Asp 180 Leu Leu Val His Trp Cys His Gly Ala Pro Gly Val Ile Tyr Met Leu 200 195 Ile Gln Ala Tyr Lys Val Phe Arg Glu Glu Lys Tyr Leu Cys Asp Ala 220 210 ... 215 Tyr Gln Cys Ala Asp Val Ile Trp Gln Tyr Gly Leu Leu Lys Lys Gly 230

PCT/IB98/02122 -

Tyr Gly Leu Cys His Gly Ser Ala Gly Asn Ala Tyr Ala Phe Leu Thr
240

Leu Tyr Asn Leu Thr Gln Asp Met Lys Tyr Leu Tyr Arg Ala Cys Lys
255

260

Phe Ala Glu Trp Cys Leu Glu Tyr Gly Glu His Gly Cys Arg Thr Pro
275

Asp Thr Pro Phe Ser Leu Phe Glu Gly Met Ala Gly Thr Ile Tyr Phe
290

Leu Ala Asp Leu Leu Val Pro Thr Lys Ala Arg Phe Pro Ala Phe Glu
305

Leu His Gly Cys Arg Thr Pro
285

Asp Thr Pro Phe Ser Leu Phe Glu Gly Met Ala Gly Thr Ile Tyr Phe
296

Leu Ala Asp Leu Leu Val Pro Thr Lys Ala Arg Phe Pro Ala Phe Glu
315

<210> 197 <211> 30 <212> PRT <213> Homo sapiens

<210> 198 <211> 112 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -48..-1

<400> 198 Met Gln Asp Thr Gly Ser Val Val Pro Leu His Trp Phe Gly Phe Gly -40 -45 Tyr Ala Ala Leu Val Ala Ser Gly Gly Ile Ile Gly Tyr Val Lys Ala -25 -20 -30 Gly Ser Val Pro Ser Leu Ala Ala Gly Leu Leu Phe Gly Ser Leu Ala -10 - 5 Gly Leu Gly Ala Tyr Gln Leu Ser Gln Asp Pro Arg Asn Val Trp Val 10 Phe Leu Ala Thr Ser Gly Thr Leu Ala Gly Ile Met Gly Met Arg Phe 25 Tyr His Ser Gly Lys Phe Met Pro Ala Gly Leu Ile Ala Gly Ala Ser

35 40 45
Leu Leu Met Val Ala Lys Val Gly Val Ser Met Phe Asn Arg Pro His
50 55 60

<210> 199 <211> 54 <212> PRT <213> Homo sapiens

 Pro
 Arg
 Trp
 His
 Arg
 Leu
 Pro
 Pro
 Gln
 Ser
 Leu
 Gln
 His
 His
 Gln
 Tyr

 Cys
 Gln
 Arg
 Arg
 Trp
 Pro
 Asp
 Arg
 Cys
 Leu
 Gln
 Ser
 His
 Thr
 Gln

 Ser
 Ser
 Gly
 His
 Leu
 Pro
 Fro
 F

<210> 200 <211> 151 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21...1

<400> 200 Met Ala Ala Ser Thr Ser Met Xaa Pro Val Ala Val Thr Ala Ala Val -15 -10 Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu Arg Glu Ile Lys Lys Gln Leu Leu Ile Ala Gly Leu Thr Arg Glu Arg Gly Leu 20 Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser Leu Pro Ala 35 Leu Pro Xaa Gly Gln Leu Gln Pro Pro Pro Pro Ile Thr Glu Glu Asp . .55 5.0 Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr Phe Asp Val 70 65 Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys Asn Ser Lys 85 Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val Arg Ala Ile 100 Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe Arg Thr Asn 115 110 Gly Lys Val Lys Ser Phe Lys

<210> 201 <211> 228 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -25..-1

Leu Met Ile Thr Ala Ile Leu Leu Gly Phe Leu Gly Leu Leu Leu Gly 60 65 Ile Ala Gly Leu Arg Cys Thr Asn Ile Gly Gly Leu Glu Leu Ser Arg 80 75 Lys Ala Lys Leu Ala Ala Thr Ala Gly Ala Pro His Ile Leu Ala Gly 90 Ile Cys Gly Met Val Ala Ile Ser Trp Tyr Ala Phe Asn Ile Thr Arg 115 110 Asp Phe Phe Asp Pro Leu Tyr Pro Gly Thr Lys Tyr Glu Leu Gly Pro 130 125 Ala Leu Tyr Leu Gly Trp Ser Ala Ser Leu Ile Ser Ile Leu Gly Gly 145 140 Leu Cys Leu Cys Ser Ala Cys Cys Cys Gly Ser Asp Glu Asp Pro Ala 160 155 Ala Ser Ala Arg Arg Pro Tyr Gln Ala Pro Val Ser Val Met Pro Val 170 175 Ala Thr Ser Asp Gln Glu Gly Asp Ser Ser Phe Gly Lys Tyr Gly Arg 190 Asn Ala Tyr Val 200

<210 > 202 <211 > 64 <212 > PRT <213 > Homo sapiens <220 > <221 > SIGNAL <222 > -47..-1

<210> 203
<211> 146
<212> PRT
<213> Homo sapiens

<220>
<221> SIGNAL
<222> -31..-1

<400> 203

<210> 204 <211> 87 <212> PRT <213> Homo sapiens

<210> 205 <211> 40 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1

<210> 206 <211> 154 <212> PRT <213> Homo sapiens

<400> 206
Met Gly Ser Leu Ser Gly Leu Arg Leu Ala Ala Gly Ser Cys Phe Arg

10 Leu Cys Glu Arg Asp Val Ser Ser Ser Leu Arg Leu Thr Arg Ser Ser 25 20 Asp Leu Lys Arg Ile Asn Gly Phe Cys Thr Lys Pro Gln Glu Ser Pro 40 Gly Ala Pro Ser Arg Thr Tyr Asn Arg Val Pro Leu His Lys Pro Thr 55 Asp Trp Gln Lys Lys Ile Leu Ile Trp Ser Gly Arg Phe Lys Lys Glu 75 70 Asp Glu Ile Pro Glu Thr Val Ser Leu Glu Met Leu Asp Ala Ala Lys 90 85 Asn Lys Met Arg Val Lys Ser Ser Tyr Leu Met Ile Ala Leu Thr Val 105 Val Gly Cys Ile Phe Met Val Ile Glu Gly Lys Lys Ala Ala Gln Arg 125 120 His Glu Thr Leu Thr Ser Leu Asn Leu Glu Lys Lys Ala Arg Leu Lys 140 135 Glu Glu Ala Ala Met Lys Ala Lys Thr Glu 150

<210> 207 <211> 101 <212> PRT <213> Homo sapiens

<210> 208
<211> 456
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -22..-1

```
30
                               35
Glu Glu Glu Glu Glu Arg Lys Lys Cys Pro Lys Lys Ala Ser
                                      55
                          50
Phe Ala Ser Ala Ser Ala Glu Val Gly Lys Lys Gly Lys Lys Lys Cys
                       65
                                          70
Gln Lys Gln Gly Pro Pro Cys Ser Asp Ser Glu Glu Glu Val Glu Arg
                   80
                                      85
Lys Lys Lys Cys His Lys Gln Ala Leu Val Gly Ser Asp Ser Ala Glu
              95
                                  100
Asp Glu Lys Arg Lys Arg Lys Cys Gln Lys His Ala Pro Ile Asn Ser
           110
                              115
Ala Gln His Leu Asp Asn Val Asp Gln Thr Gly Pro Lys Ala Trp Lys
                           130
       125
                                             135
Gly Ser Thr Thr Asn Asp Pro Pro Lys Gln Ser Pro Gly Ser Thr Ser
                                          150
                      145
Pro Lys Pro Pro His Thr Leu Ser Arg Lys Gln Trp Arg Asn Arg Gln
                   160
                                      165
Lys Asn Lys Arg Arg Cys Lys Asn Lys Phe Gln Pro Pro Gln Val Pro
               175
                                  180
Asp Gln Ala Pro Ala Glu Ala Pro Thr Glu Lys Thr Glu Val Ser Pro
           190
                               195
Val Pro Arg Thr Asp Ser His Gly Ala Arg Ala Gly Ala Leu Arg Ala
                           210
Arg Met Ala Gln Arg Leu Asp Gly Ala Arg Phe Arg Tyr Leu Asn Glu
                       225
                                          230
Gln Leu Tyr Ser Gly Pro Ser Ser Ala Ala Gln Arg Leu Phe Gln Glu
                   240
                                       245
Asp Pro Glu Ala Phe Leu Leu Tyr His Arg Gly Phe Gln Ser Gln Val
               255
                                  260
Lys Lys Trp Pro Leu Gln Pro Val Asp Arg Ile Ala Arg Asp Leu Arg
           270
                               275
                                                  280
Gln Arg Pro Ala Ser Leu Val Val Ala Asp Phe Gly Cys Gly Asp Cys
                          290
                                              295
Arg Leu Ala Ser Ser Ile Arg Asn Pro Val His Cys Phe Asp Leu Ala
                       305
                                          310
Ser Leu Asp Pro Arg Val Thr Val Cys Asp Met Ala Gln Val Pro Leu
                   320
                                      325
Glu Asp Glu Ser Val Asp Val Ala Val Phe Cys Leu Ser Leu Met Gly
                                  340
Thr Asn Ile Arg Asp Phe Leu Glu Glu Ala Asn Arg Val Leu Lys Pro
                               355
Gly Gly Leu Leu Lys Val Ala Glu Val Ser Ser Arg Phe Glu Asp Val
                           370
Arg Thr Phe Leu Arg Ala Val Thr Lys Leu Gly Phe Lys Ile Val Ser
                      385
Lys Asp Leu Thr Asn Ser His Phe Phe Leu Phe Asp Phe Gln Lys Thr
                  400
                                      405
Gly Pro Pro Leu Val Gly Pro Lys Ala Gln Leu Ser Gly Leu Gln Leu
              415
                                 420
Gln Pro Cys Leu Tyr Lys Arg Arg
           430
```

<210> 209

<211> 98

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -17..-1

```
<400> 209
Met Pro Ser Ser Phe Phe Leu Leu Leu Gln Phe Phe Leu Arg Ile Asp
             -10
    -15.
Gly Val Leu Ile Arg Met Asn Asp Thr Arg Leu Tyr His Glu Ala Asp
                                     10
Lys Thr Tyr Met Leu Arg Glu Tyr Thr Ser Arg Glu Ser Lys Ile Ser
                                 25
Ser Leu Met His Val Pro Pro Ser Leu Phe Thr Glu Pro Asn Glu Ile
                        . 40
Ser Gln Tyr Leu Pro Ile Lys Glu Ala Val Cys Glu Lys Leu Ile Phe
                   55
Pro Glu Arg Ile Asp Pro Asn Pro Ala Asp Ser Gln Lys Ser Thr Gln
                   70
Val Glu
80
<210> 210
<211> 83
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -29..-1
<400> 210
Met Thr Leu Leu Ser Phe Ala Ala Phe Thr Ala Ala Phe Ser Val Leu
                                  -20
               -25
Pro Cys Tyr Tyr Leu Gly Leu Phe Gln Arg Ala Leu Ala Ser Val Phe
                        -5
           -10
Asp Pro Leu Cys Val Cys Ser Arg Val Leu Pro Thr Pro Val Cys Thr
                      10
Leu Val Ala Thr Gln Ala Glu Lys Ile Leu Glu Asn Gly Pro Cys Pro
                  25
                                     30
Thr Lys Glu Ala Ala Gln Leu Val Gly Lys Gly Ser Val Ser Ala Arg
                                  45
Asn Ala Ser
<210> 211
<211> 229
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -23..-1
<400> 211
Met Gly Asp Lys Ile Trp Leu Pro Phe Pro Val Leu Leu Ala Ala
                               -15
            -20
Leu Pro Pro Val Leu Leu Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu
     -5
Asp Ser Asp Phe Thr Phe Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe
                                       20
                  15
Tyr Gln Pro Met Pro Leu Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val
                30
                                   35
Leu Asp Gly Ala Gly Leu Asp Ile Asp Phe His Leu Ala Ser Pro Glu
```

```
50
Gly Lys Thr Leu Val Phe Glu Gln Arg Lys Ser Asp Gly Val His Thr
                          65
Val Glu Thr Glu Val Gly Asp Tyr Met Phe Cys Phe Asp Asn Thr Phe
                                         85
Ser Thr Ile Ser Glu Lys Val Ile Phe Phe Glu Leu Ile Leu Asp Asn
                                     100
Met Gly Glu Gln Ala Gln Glu Glu Asp Trp Lys Lys Tyr Ile Thr
                                 115
               110
Gly Thr Asp Ile Leu Asp Met Lys Leu Glu Asp Ile Leu Glu Ser Ile
                             130
           125
Ser Ser Ile Lys Ser Arg Leu Ser Lys Ser Gly His Ile Gln Ile Leu
                         145
Leu Arg Ala Phe Glu Ala Arg Asp Arg Asn Ile Gln Glu Ser Asn Phe
                                        165
                    160
Asp Arg Val Asn Phe Trp Ser Met Val Asn Leu Val Val Met Val Val
                              180
            175
Val Ser Ala Ile Gln Val Tyr Met Leu Lys Ser Leu Phe Glu Asp Lys
                          195
              190
Arg Lys Ser Arg Thr
           205
```

<210> 212 <211> 152 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1

<400> 212 Met Ala Gln Leu Gly Ala Val Val Ala Val Ala Ser Ser Phe Phe Cys -15 Ala Ser Leu Phe Ser Ala Val His Lys Ile Glu Glu Gly His Ile Gly Val Tyr Tyr Arg Gly Gly Ala Leu Leu Thr Ser Thr Ser Gly Pro Gly 20 15 Phe His Leu Met Leu Pro Phe Ile Thr Ser Tyr Lys Ser Val Gln Thr Thr Leu Gln Thr Asp Glu Val Lys Asn Val Pro Cys Gly Thr Ser Gly Gly Val Met Ile Tyr Phe Asp Arg Ile Glu Val Val Asn Phe Leu Val 70 Pro Asn Ala Val His Asp Ile Val Lys Asn Tyr Thr Ala Asp Tyr Asp 85 Lys Ala Leu Ile Phe Asn Lys Ile His His Glu Leu Asn Gln Phe Cys 100 Ser Val His Thr Leu Gln Glu Val Tyr Ile Glu Leu Phe Gly Leu Glu 115 Asn Asp Phe Ser Gln Glu Ser Ser 130

<210> 213 <211> 179 <212> PRT <213> Home sapiens

```
<220>
<221> SIGNAL
<222> -54..-1
```

<400> 213

Met Ala Ala Ser Glu Ala Ala Val Val Ser Ser Pro Ser Leu Lys Thr -50 -45 Asp Thr Ser Pro Val Leu Glu Thr Ala Gly Thr Val Ala Ala Met Ala -35 -30 -25 Ala Thr Pro Ser Ala Arg Ala Ala Ala Ala Val Val Ala Ala Ala Ala -20 -15 -10 Arg Thr Gly Ser Glu Ala Arg Val Ser Lys Ala Ala Leu Ala Thr Lys Leu Leu Ser Leu Ser Gly Val Phe Ala Val His Lys Pro Lys Gly Pro 20 15 Thr Ser Ala Glu Leu Leu Asn Arg Leu Lys Glu Lys Leu Leu Ala Glu 30 35 Ala Gly Met Pro Ser Pro Glu Trp Thr Lys Arg Lys Lys Gln Thr Leu 50 Lys Ile Gly His Gly Gly Thr Leu Asp Ser Ala Ala Arg Gly Val Leu Val Val Gly Ile Gly Ser Gly Thr Lys Met Leu Thr Ser Met Leu Ser 80 Gly Ser Lys Arg Tyr Thr Ala Ile Gly Glu Leu Gly Lys Ala Thr Asp 95 100 Thr Leu Asp Ser Thr Gly Lys Val Thr Glu Glu Lys Pro Tyr Gly Met 115 110 Asn Leu Ile

<210> 214 <211> 269 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -92..-1

125

<400> 214 Met Ile Thr His Val Thr Leu Glu Asp Ala Leu Ser Asn Val Asp Leu -85 Leu Glu Glu Leu Pro Leu Pro Asp Gln Gln Pro Cys Ile Glu Pro Pro -70 -65 Pro Ser Ser Ile Met Tyr Gln Ala Asn Phe Asp Thr Asn Phe Glu Asp -55 - -50 Arg Asn Ala Phe Val Thr Gly Ile Ala Arg Tyr Ile Glu Gln Ala Thr -40 -35 Val His Ser Ser Met Asn Glu Met Leu Glu Glu Gly His Glu Tyr Ala -25 -20 Val Met Leu Tyr Thr Trp Arg Ser Cys Ser Arg Ala Ile Pro Gln Val - 5 Lys Cys Asn Glu Gln Pro Asn Arg Val Glu Ile Tyr Glu Lys Thr Val 10 15 Glu Val Leu Glu Pro Glu Val Thr Lys Leu Met Lys Phe Met Tyr Phe 30 Gln Arg Lys Ala Ile Glu Arg Phe Cys Ser Glu Val Lys Arg Leu Cys 45 His Ala Glu Arg Arg Lys Asp Phe Val Ser Glu Ala Tyr Leu Leu Thr 60 65

Leu Gly Lys Phe Ile Asn Met Phe Ala Val Leu Asp Glu Leu Lys Asn 75 Met Lys Cys Ser Val Lys Asn Asp His Ser Ala Tyr Lys Arg Ala Ala 90 95 Gln Phe Leu Arg Lys Met Ala Asp Pro Gln Ser Ile Gln Glu Ser Gln 105 110 Asn Leu Ser Met Phe Leu Ala Asn His Asn Arg Ile Thr Gln Cys Leu 120 125 His Gln Gln Leu Glu Val Ile Pro Gly Tyr Glu Glu Leu Leu Ala Asp 140 145 Ile Val Asn Ile Cys Val Asp Tyr Tyr Glu Asn Lys Met Tyr Leu Thr 155 Pro Ser Glu Lys His Met Leu Leu Lys Val Lys Leu Pro 165 170

```
<210> 215
<211> 135
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -22..-1
<400> 215
Met Gln Thr Val Tyr Tyr Gly Ser Leu Gly Leu Trp Leu Ala Leu Val
                           -15
Asp Gly Leu Val Arg Ser Ser Pro Ser Leu Asp Gln Met Phe Asp Ala
                      1
                                    5
Glu Ile Leu Gly Phe Ser Thr Pro Pro Gly Arg Leu Ser Met Met Ser
               15
                                   20
Phe Ile Phe Asn Ala Leu Thr Cys Ala Leu Gly Leu Leu Tyr Phe Ile
                              35
                                                  40
Arg Arg Gly Lys Gln Cys Leu Asp Phe Thr Val Thr Val His Phe Phe
                           50
His Leu Leu Gly Cys Trp Phe Tyr Ser Ser Arg Phe Pro Ser Ala Leu
                       65
                                           70
Thr Trp Trp Leu Val Gln Ala Val Cys Ile Ala Leu Met Ala Val Ile
                                       85
Gly Glu Tyr Leu Cys Met Arg Thr Glu Leu Lys Glu Ile Pro Leu Asn
               95
                                   100
Ser Ala Pro Lys Ser Asn Val
```

<210> 216 <211> 67 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -38..-1 <400> 216 Met Asn Asn Val Gln Pro Lys

110

Met Asn Asn Val Gln Pro Lys Ile Lys His Arg Pro Phe Cys Phe Ser
-35 -30 -25

Val Lys Gly His Val Lys Met Leu Arg Leu Val Phe Ala Leu Val Thr
-20 -15 -10

<210> 217 <211> 125 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -54..-1 <400> 217 Met Ala Asp Glu Glu Leu Glu Ala Leu Arg Arg Gln Arg Leu Ala Glu -50 -45 Leu Gln Ala Lys His Gly Asp Pro Gly Asp Ala Ala Gln Gln Glu Ala -35 -30 Lys His Arg Glu Ala Glu Met Arg Asn Ser Ile Leu Ala Gln Val Leu -20 -15 Asp Gln Ser Ala Arg Ala Arg Leu Ser Asn Leu Ala Leu Val Lys Pro Glu Lys Thr Lys Ala Val Glu Asn Tyr Leu Ile Gln Met Ala Arg Tyr 15 20 Gly Gln Leu Ser Glu Lys Val Ser Glu Gln Gly Leu Ile Glu Ile Leu Lys Lys Val Ser Gln Gln Thr Glu Lys Thr Thr Thr Val Lys Phe Asn 50 Arg Arg Lys Val Met Asp Ser Asp Glu Asp Asp Asp Tyr 60

<210> 218 <211> 376 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1

<400> 218

 Met
 Gly
 His
 Arg
 Phe
 Leu
 Arg
 Gly
 Leu
 Leu
 Thr
 Leu
 Pro
 Glu
 Ser
 Val
 Pro

 Pro
 Pro
 Leu
 Thr
 Arg
 Ser
 Arg
 Ser
 Leu
 Met
 Ala
 Pro
 Pro
 Arg
 Ile
 Gly

 Pro
 Pro
 Lys
 Arg
 Ser
 Arg
 Ser
 Leu
 Met
 Ala
 Pro
 Pro
 Arg
 Ile
 Gly

 Pro
 Pro
 Lys
 Arg
 Ser
 Lys
 Leu
 Met
 Ala
 Pro
 Pro
 Arg
 Ile
 Gly
 Arg
 Ile
 Gly
 Arg
 Ile
 Arg
 Ile
 Arg
 Ile
 Ile

```
100
Leu Ser Ser Ala Gly Leu Ile Tyr Leu His Phe Gly His Lys Leu Leu
                                              120
                           115
Ala Gln Leu Leu Gly Thr Ser Glu Glu Asp Ser Met Val Gly Thr Leu
                                          135
                       130
Tyr Asp Lys Met Tyr Glu Asn Phe Val Glu Glu Val Asp Ala Val Asp
                                       150
                   145
Asn Gly Ile Ser Gln Trp Ala Glu Gly Glu Pro Arg Tyr Ala Leu Thr
                                   165
               160
Thr Thr Leu Ser Ala Arg Val Ala Arg Leu Asn Pro Thr Trp Asn His
                               180
           175
Pro Asp Gln Asp Thr Glu Ala Gly Phe Lys Arg Ala Met Asp Leu Val
                                               200
                           195
Gln Glu Glu Phe Leu Gln Arg Leu Asp Phe Tyr Gln His Ser Trp Leu
                        210
Pro Ala Arg Ala Leu Val Glu Glu Ala Leu Ala Gln Arg Phe Gln Val
                                       230
                    225
Asp Pro Ser Gly Glu Ile Val Glu Leu Ala Lys Gly Ala Cys Pro Trp
                                   245
                240
Lys Glu His Leu Tyr His Leu Glu Ser Gly Leu Ser Pro Pro Val Ala
                               260
Ile Phe Phe Val Ile Tyr Thr Asp Gln Ala Gly Gln Trp Arg Ile Gln
                                           280
                           275
Cys Val Pro Lys Glu Pro His Ser Phe Gln Ser Arg Leu Pro Leu Pro
                                          295
                       290
Glu Pro Trp Arg Gly Leu Arg Asp Glu Ala Leu Asp Gln Val Ser Gly
                                       310
                    305
 Ile Pro Gly Cys Ile Phe Val His Ala Ser Gly Phe Ile Gly Gly His
                                   325
                320
 Arg Thr Arg Glu Gly Ala Leu Ser Met Ala Arg Ala Thr Leu Ala Gln
                               340
            335
 Arg Ser Tyr Leu Pro Gln Ile Ser
        350
```

<210> 219 <211> 211 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1

 <400> 219

 Met Gly Glu Ala Ser - 25
 Pro Pro Ala Pro Ala Arg Arg His Leu Leu Leu -15

 Leu Leu Leu Leu Leu Leu Ser Thr Leu Val Ile Pro Ser Ala Ala Ala Pro -10

 Ile His Asp Ala Asp Ala Gln Glu Ser Ser Leu Gly Leu Thr Gly Leu 5

 Gln Ser Leu Leu Gln Gly Phe Ser Arg Leu Phe Leu Lys Gly Asn Leu 20

 Leu Arg Gly Ile Asp Ser Leu Phe Ser Ala Pro Met Asp Phe Arg Gly 35

 Leu Pro Gly Asn Tyr His Lys Glu Glu Asn Gln Glu His Gln Leu Gly 55

 Asn Asn Thr Leu Ser Ser His Leu Gln Ile Asp Lys Val Pro Arg Met 70

 Glu Glu Lys Glu Ala Leu Val Pro Ile Gln Lys Ala Thr Asp Ser Phe 85

His Thr Glu Leu His Pro Arg Val Ala Phe Trp Ile Ile Lys Leu Pro 105 110 Arg Arg Ser His Gln Asp Ala Leu Glu Gly Gly His Trp Leu Ser 120 125 Glu Lys Arg His Arg Leu Gln Ala Ile Arg Asp Gly Leu Arg Lys Gly 135 140 Thr His Lys Asp Val Leu Glu Glu Gly Thr Glu Ser Ser Ser His Ser 150 155 Arg Leu Ser Pro Arg Lys Thr His Leu Leu Tyr Ile Leu Arg Pro Ser 170 Arg Gln Leu 180

<210> 220 <211> 154 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -60..-1 <400> 220 Met Gly Ser Lys Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu -55 Arg Gln Arg Arg Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys -40 -35 Arg Val Lys Ala Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu -25 -20 -15 Val Arg Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln -10 -5 Cys Trp Trp Arg Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln 10 15 Ala Leu Leu Arg Val Tyr Val Ile Gln Glu Gln Ala Thr Val Lys Leu 30 Gln Ser Cys Ile Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met 45 Cys Asn Ala Leu Cys Leu Phe Gln Val Pro Glu Ser Ser Leu Ala Phe 60 Gln Thr Asp Gly Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln 75 Pro Glu Phe His Ile Glu Ile Leu Ser Ile

<210> 221 <211> 123 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -42..-1

Ala Val Ser Leu Ser Ala Pro Ala Phe Ala Ser Ala Leu Arg Ser Met
-10
-5
Lys Ser Ser Gln Ala Ala Arg Lys Asp Asp Phe Leu Arg Ser Leu Ser
10
Asp Gly Asp Ser Gly Thr Ser Glu His Ile Ser Ala Val Val Thr Ser
25
Pro Arg Ile Ser Cys His Gly Ala Ala Ile Pro Thr Ala Arg Ala Leu
40
Cys Leu Gly Cys Ser Cys Cys Thr Glu Arg Leu Leu Pro Pro
55
Ser Leu Leu Ser Leu Glu Ala Pro Ala Ser Thr
75

<210> 222 <211> 346 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 222 Met Ala Met Ala Gln Lys Leu Ser His Leu Leu Pro Ser Leu Arg Gln -10 -15 Val Ile Gln Glu Pro Gln Leu Ser Leu Gln Pro Glu Pro Val Phe Thr Val Asp Arg Ala Glu Val Pro Pro Leu Phe Trp Lys Pro Tyr Ile Tyr 25 20 Ala Gly Tyr Arg Pro Leu His Gln Thr Trp Arg Phe Tyr Phe Arg Thr 40 35 Leu Phe Gln Gln His Asn Glu Ala Val Asn Val Trp Thr His Leu Leu 55 50 Ala Ala Leu Val Leu Leu Leu Arg Leu Ala Leu Phe Val Glu Thr Val 70 Asp Phe Trp Gly Asp Pro His Ala Leu Pro Leu Phe Ile Ile Val Leu 85 Ala Ser Phe Thr Tyr Leu Ser Leu Ser Ala Leu Ala His Leu Leu Gln 100 Ala Lys Ser Glu Phe Trp His Tyr Ser Phe Phe Phe Leu Asp Tyr Val 120 115 Gly Val Ala Val Tyr Gln Phe Gly Ser Ala Leu Ala His Phe Tyr Tyr 135 130 Ala Ile Glu Pro Ala Trp His Ala Gln Val Gln Ala Val Phe Leu Pro 145 150 Met Ala Ala Phe Leu Ala Trp Leu Ser Cys Ile Gly Ser Cys Tyr Asn Lys Tyr Ile Gln Lys Pro Gly Leu Leu Gly Arg Thr Cys Gln Glu Val 180 185 Pro Ser Val Leu Ala Tyr Ala Leu Asp Ile Ser Pro Val Val His Arg 200 195 Ile Phe Val Ser Ser Asp Pro Thr Thr Asp Asp Pro Ala Leu Leu Tyr 215 210 His Lys Cys Gln Val Val Phe Phe Leu Leu Ala Ala Phe Phe Ser 230 Thr Phe Met Pro Glu Arg Trp Phe Pro Gly Ser Cys His Val Phe Gly 245 Gln Gly His Gln Leu Phe His Ile Phe Leu Val Leu Cys Thr Leu Ala 260 Gln Leu Glu Ala Val Ala Leu Asp Tyr Glu Ala Arg Arg Pro Ile Tyr

```
<210> 223
<211> 210
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -20..-1
<400> 223
Met Asp Asn Arg Phe Ala Thr Ala Phe Val Ile Ala Cys Val Leu Ser
-20
                                     -10
               -15
Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp
               1
Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys
       15
                          20
Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr
                                          40
Asn Asp Ala Leu Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Arg Arg
                   50
                                      55
Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg
                                  70
Thr Glu Ser Phe Asp Val Val Thr Lys Cys Val Ser Phe Thr Leu Thr
                              85
Glu Gln Phe Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly
                          100
                                             105
Ile Asp Leu Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro
                       115
                                         120
Phe Val Ser Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys
                                    135
                   130
Ala Cys Ile Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu
                       150
His Leu Leu Ala Val Thr Lys Glu Ser Met Leu Pro Ala Gly Ala Glu
                 165
Ser Lys His Thr Ala Thr Pro Ala His Ala Cys Val Gln Thr Gly Lys
             180
Pro Lys
  190
```

```
<210> 224
<211> 184
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -20..-1
```

<400> 224

Met Asp Asn Arg Phe Ala Thr Ala Phe Val Ile Ala Cys Val Leu Ser

```
-10
                  -15
Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp
                          5 .
Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys
                         20
Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr
                  35
Asn Asp Ala Pro Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Arg Arg
                 50
Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg
                                70
Thr Glu Ser Phe Asp Val Val Thr Lys Cys Val Ser Phe Thr Leu Thr
          80
                            85
Glu Gln Phe Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly
                        100
Ile Asp Leu Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro
                  115
                                       120
Phe Val Ser Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys
                          135
                130
Ala Cys Ile Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu
                       150
            145
His Leu Leu Ala Asp Thr Met Leu
          160
```

```
<210> 225
<211> 227
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -22..-1
<400> 225
Met Gly Trp Thr Met Arg Leu Val Thr Ala Ala Leu Leu Leu Gly Leu
                   -15
Met Met Val Val Thr Gly Asp Glu Asp Glu Asn Ser Pro Cys Ala His
Glu Ala Leu Leu Asp Glu Asp Thr Leu Phe Cys Gln Gly Leu Glu Val
Phe Tyr Pro Glu Leu Gly Asn Ile Gly Cys Lys Val Val Pro Asp Cys
                              35
Asn Asn Tyr Arg Gln Lys Ile Thr Ser Trp Met Glu Pro Ile Val Lys
                          50
Phe Pro Gly Ala Val Asp Gly Ala Thr Tyr Ile Leu Val Met Val Asp
                      65
Pro Asp Ala Pro Ser Arg Ala Glu Pro Arg Gln Arg Phe Trp Arg His
                                    85
                - 80
Trp Leu Val Thr Asp Ile Lys Gly Ala Asp Leu Lys Lys Gly Lys Ile
                                 100
Gln Gly Gln Glu Leu Ser Ala Tyr Gln Ala Pro Ser Pro Pro Ala His
                             115
          110
Ser Gly Phe His Arg Tyr Gln Phe Phe Val Tyr Leu Gln Glu Gly Lys
                         130
Val Ile Ser Leu Leu Pro Lys Glu Asn Lys Thr Arg Gly Ser Trp Lys
                              150
               145
Met Asp Arg Phe Leu Asn Arg Phe His Leu Gly Glu Pro Glu Ala Ser
               160 165
Thr Gln Phe Met Thr Gln Asn Tyr Gln Asp Ser Pro Thr Leu Gln Ala
                                 180
```

Pro Arg Glu Arg Ala Ser Glu Pro Lys His Lys Asn Gln Ala Glu Ile
190 195 200

Ala Ala Cys
205

<210> 226 <211> 74 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -41..-1

<400> 226

 Met
 Ile
 Ala
 Arg
 Arg
 Asn
 Pro
 Val
 Pro
 Leu
 Arg
 Pro
 Asp
 Glu

 -40
 -40
 -80
 -35
 -80
 -30
 -30
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -80
 -15
 -15
 -15
 -80
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -10
 -

<210> 227 <211> 73 <212> PRT <213> Homo sapiens <400> 227 Met Glu Lys Tyr Glu Asn Leu Gly Leu Val Gly Glu Gly Ser Tyr Gly 15 10 5 Met Val Met Lys Cys Arg Asn Lys Asp Thr Gly Arg Ile Val Ala Ile 20 25 Lys Lys Phe Leu Glu Ser Asp Asp Lys Met Val Lys Lys Ile Ala 40 Met Arg Glu Val Lys Leu Leu Lys Gln Leu Arg His Glu Asn Leu Val Asn Leu Leu Glu Val Cys Lys Lys 70

<210> 228
<211> 82
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1

<210> 229 <211> 119 <212> PRT <213> Homo sapiens <220>

<221> SIGNAL <222> -56..-1

<400> 229 Met Ala Glu Pro Ser Ala Ala Thr Gln Ser His Ser Ile Ser Ser Ser -45 -50 Ser Phe Gly Ala Glu Pro Ser Ala Pro Gly Gly Gly Ser Pro Gly -35 -30 Ala Cys Pro Ala Leu Gly Thr Lys Ser Cys Ser Ser Ser Cys Ala Asp -15 -20 Ser Phe Val Ser Ser Ser Ser Gln Pro Val Ser Leu Phe Ser Thr 5 -5 Ser Gln Glu Gly Leu Ser Ser Leu Cys Ser Asp Glu Pro Ser Ser Glu 15 Ile Met Thr Ser Ser Phe Leu Ser Ser Ser Glu Ile His Asn Thr Gly 35 Leu Thr Ile Leu His Gly Glu Lys Ser His Val Leu Gly Ser Gln Pro 50 Ile Leu Ala Lys Lys Lys Lys

<210> 230 <211> 54 <212> PRT <213> Homo sapiens

60

<210> 231 <211> 210 <212> PRT <213> Homo sapiens <221> SIGNAL <222> -14..-1 <400> 231 Met Leu Thr Leu Leu Gly Leu Ser Phe Ile Leu Ala Gly Leu Ile Val -5 -10 Gly Gly Ala Cys Ile Tyr Lys Tyr Phe Met Pro Lys Ser Thr Ile Tyr 10 Arg Gly Glu Met Cys Phe Phe Asp Ser Glu Asp Pro Ala Asn Ser Leu 25 Arg Gly Gly Glu Pro Asn Phe Leu Pro Val Thr Glu Glu Ala Asp Ile 40 Arg Glu Asp Asp Asn Ile Ala Ile Ile Asp Val Pro Val Pro Ser Phe Ser Asp Ser Asp Pro Ala Ala Ile Ile His Asp Phe Glu Lys Gly Met Thr Ala Tyr Leu Asp Leu Leu Leu Gly Ile Cys Tyr Leu Met Pro Leu 90 Asn Thr Ser Ile Val Met Pro Pro Lys Asn Leu Val Glu Leu Phe Gly 105 Lys Leu Ala Ser Gly Arg Tyr Leu Pro Gln Thr Tyr Val Val Arg Glu 120 Asp Leu Val Ala Val Glu Glu Ile Arg Asp Val Ser Asn Leu Gly Ile 140 135 Phe Ile Tyr Gln Leu Cys Asn Asn Arg Lys Ser Phe Arg Leu Arg Arg 155 150 Arg Asp Leu Leu Gly Phe Asn Lys Arg Ala Ile Asp Lys Cys Trp 175 170 Lys Ile Arg His Phe Pro Asn Glu Phe Ile Val Glu Thr Lys Ile Cys 185 Gln Glu

<210> 232 <211> 108 <212> PRT <213> Homo sapiens

<220>

```
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -18..-1
<400> 233
Met Ser Ser Gly Arg Leu Arg Trp Leu Met Pro Val Ile Pro Ala Leu
       -15
                   -10
Trp Gly Ala Glu Lys Gly Glu Ser Pro Glu Val Ser Ser Phe Glu Thr
                     . 5
Arg Leu Ala Asn Met Ala Lys Pro Cys Leu Tyr
       20
<210> 234
<211> 36
<212> PRT
<213> Homo sapiens
<400> 234
Met Ser Ala Arg Ile Pro Phe Tyr Lys Asp Thr Ser Gln Ile Arg Leu
                                 10
Gly Ser Thr Ile Ile Pro His Phe Asn Leu Ile Thr Phe Val Lys Thr
    20
                           25
Phe Phe Gln Ile
   35
<210> 235
<211> 307
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -13..-1
<400> 235
Met Leu Ala Val Ser Leu Thr Val Pro Leu Leu Gly Ala Met Met Leu
                          - 5
           -10
Leu Glu Ser Pro Ile Asp Pro Gln Pro Leu Ser Phe Lys Glu Pro Pro
                      10
                                         15
Leu Leu Gly Val Leu His Pro Asn Thr Lys Leu Arg Gln Ala Glu
                   25
                                      30
Arg Leu Phe Glu Asn Gln Leu Val Gly Pro Glu Ser Ile Ala His Ile
                                  45
Gly Asp Val Met Phe Thr Gly Thr Ala Asp Gly Arg Val Val Lys Leu
                              60
Glu Asn Gly Glu Ile Glu Thr Ile Ala Arg Phe Gly Ser Gly Pro Cys
                          75
Lys Thr Arg Asp Asp Glu Pro Val Cys Gly Arg Pro Leu Gly Ile Arg
                      90
Ala Gly Pro Asn Gly Thr Leu Phe Val Ala Asp Ala Cys Lys Gly Leu
                  105
                                      110
Phe Glu Val Asn Pro Trp Lys Arg Glu Val Lys Leu Leu Ser Ser
                                 125
              120
```

Glu Thr Pro Ile Glu Gly Lys Asn Met Ser Phe Val Asn Asp Leu Thr 135 140 145 Val Ser Gln Asp Gly Arg Lys Ile Tyr Phe Thr Asp Ser Ser Ser Lys 155 Trp Gln Arg Arg Asp Tyr Leu Leu Val Met Glu Gly Thr Asp Asp 170 175 Gly Arg Leu Leu Glu Tyr Asp Thr Val Thr Arg Glu Val Lys Val Leu 185 190 Leu Asp Gln Leu Arg Phe Pro Asn Gly Val Gln Leu Ser Pro Ala Glu 205 Asp Phe Val Leu Val Ala Glu Thr Thr Met Ala Arg Ile Arg Arg Val 215 220 Tyr Val Ser Gly Leu Met Lys Gly Gly Ala Asp Leu Phe Val Glu Asn 230 235 Met Pro Gly Phe Pro Asp Asn Ile Arg Pro Ser Ser Gly Gly Tyr 250 Trp Val Gly Met Ser Thr Ile Arg Pro Asn Pro Gly Phe Ser Met Leu 265 270 Asp Phe Leu Ser Glu Arg Pro Trp Ile Lys Arg Met Ile Phe Lys Ala 280 285 Lys Lys Lys

<211> 106 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -32..-1 <400> 236 Met Phe Ala Pro Ala Val Met Arg Ala Phe Arg Lys Asn Lys Thr Leu - 30 -25 -20 Gly Tyr Gly Val Pro Met Leu Leu Leu Ile Val Gly Gly Ser Phe Gly -10 Leu Arg Glu Phe Ser Gln Ile Arg Tyr Asp Ala Val Lys Ser Lys Met 10 Asp Pro Glu Leu Glu Lys Lys Leu Lys Glu Asn Lys Ile Ser Leu Glu 25 Ser Glu Tyr Glu Lys Ile Lys Asp Ser Lys Phe Asp Asp Trp Lys Asn 40 Ile Arg Gly Pro Arg Pro Trp Glu Asp Pro Asp Leu Leu Gln Gly Arg 55 Asn Pro Glu Ser Leu Lys Thr Lys Thr Thr

<210> 237 <211> 42 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 237

<210> 236

Met Asp Leu Arg Gln Phe Leu Met Cys Leu Ser Leu Cys Thr Ala Phe
-15 -10 -5
Ala Leu Ser Lys Pro Thr Glu Lys Lys Asp Arg Val His His Glu Pro

10 Gln Leu Ser Asp Lys Val His Asn Asp Ile 20 <210> 238 <211> 117 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -20..-1 <400> 238 Met Asp Asn Arg Phe Ala Thr Ala Phe Val Ile Ala Cys Val Leu Ser -15 -10 Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys 20 Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr 35 Asn Asp Ala Leu Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Gly Arg 50 _ . . 55 Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg 70 Thr Gly Ile Ser Leu Ile Leu Thr Ser Val Phe Phe Thr Trp Leu Ile 85 Ile Asp Lys Thr Thr 95 <210> 239 <211> 178 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1 <400> 239 Met Glu Arg Gln Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe -35 -30 -25 Gln His Xaa Xaa Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile -15 -10 Leu Thr Ile Leu Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu 20 Ile Ser Arg Tyr Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val

<210> 240 <211> 126 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1 <400> 240 Met Gln Phe Val Asn Val Gly Tyr Phe Leu Ile Ala Ala Gly Val Val -20 Val Leu Ala Leu Gly Phe Leu Gly Cys Tyr Gly Ala Lys Thr Glu Ser Met Cys Ala Leu Val Thr Phe Phe Phe Ile Leu Leu Ile Phe Ile 15 Ala Glu Val Ala Ala Ala Val Val Ala Leu Val Tyr Thr Thr Met Ala 30 Glu His Phe Leu Thr Leu Leu Val Val Pro Ala Ile Lys Lys Asp Tyr 45 50 Gly Ser Gln Glu Asp Phe Thr Gln Val Trp Asn Thr Thr Met Lys Gly 60

Leu Lys Cys Arg Gly Phe Thr Asn Tyr Thr Asp Phe Glu Asp Ser Pro

Tyr Phe Lys Met His Lys Pro Val Thr Met Lys Lys Lys

80

<210> 241 <211> 174 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -115..-1

75

<400> 241 Met Arg Trp Ser Cys Glu His Leu Val Met Val Trp Ile Asn Ala Phe -110 -105 Val Met Leu Thr Thr Gln Leu Leu Pro Ser Lys Tyr Cys Asp Leu Leu - 95 -90 His Lys Ser Ala Ala His Leu Gly Lys Trp Gln Lys Leu Glu His Gly -80 -75 Ser Tyr Ser Asn Ala Pro Gln His Ile Trp Ser Glu Asn Thr Ile Trp -60 Pro Gln Gly Val Leu Val Arg His Ser Arg Cys Leu Tyr Arg Ala Met -45 -40 Gly Pro Tyr Asn Val Ala Val Pro Ser Asp Val Ser His Ala Arg Phe -30 -25 Tyr Phe Leu Phe His Arg Pro Leu Arg Leu Leu Asn Leu Leu Ile Leu

5.0

98

193

253 313

433

553

613

853

896

<210> 242

-15 -10 -5

Ile Glu Gly Gly Val Val Phe Tyr Gln Leu Tyr Ser Leu Leu Arg Ser

1 5 10

Glu Lys Trp Asn His Thr Leu Ser Met Ala Leu Ile Leu Phe Cys Asn
15 20 25

Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Arg Ile Val Leu Gly Arg
30 35 40 45

Ala Tyr Ser Tyr Pro Leu Asn Ser Tyr Glu Leu Lys Ala Asn
50 55

<211> 896 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 18..173 <221> sig_peptide <222> 18..77 <223> Von Heijne matrix score 6.5 seq GLCVLQLTTAVTS/AF <221> polyA_signal <222> 864..869 <221> polyA_site <222> 882..893 <400> 242 aaccttcaca gtgtgag atg cct agt gtg aac agt gct gga tta tgt gtc Met Pro Ser Val Asn Ser Ala Gly Leu Cys Val -20 ttg cag ttg aca acg gca gtr acc agt gcc ttt tta cta gca aaa gtg Leu Gln Leu Thr Thr Ala Val Thr Ser Ala Phe Leu Leu Ala Lys Val 1 aat cct ttc qaa rct ttt ctc tca agg ggc ttt tgg cta tgt gcc Asn Pro Phe Glu Xaa Phe Leu Ser Arg Gly Phe Trp Leu Cys Ala Ala 15 cat cat ttc att cat cct tgc ctg gat tgagacgtgt tcctgattca His His Phe Ile His Pro Cys Leu Asp 30 aagtgttacc tcaagaagca gaagaagaaa acagactcct gatagttcag gatgcttcag agagggcagc acttatacct ggtggtcttt ctgatggtca gttttattcc cctcctgaat ccgaagcagg atctgaagaa gctgaagaaa aacaggacag tgagaaacca cttttagaac

tatgagtact acttttgtta aatgtgaaaa accctcacag aaagtcatcg aggcaaaaag

aggcaggcag tggagtctcc ctgtcgacag taaagttgaa atggtgacgt ccactgctgg ctttattgaa cagctaataa agatttattt attgtaatac ctcacagacg ttgtaccata

tccatgcaca tttagttgcc tgcctgtggc tggtaaggta atgtcatgat tcatcctctc

ttcagtgaga ctgagcctga tgtgttaaca aataggtgaa gaaagtcttg tgctgtattc ctaatcaaaa gacttaatat attgaagtaa cacttttta gtaagcaaga tacctttta tttcaattca cagaatggaa tttttttgtt tcatgtctca gatttattt gtattcttt tttaacactc tacattccc ttgttttta actcatgcac atgtgctctt tgtacagttt

taaaaagtgt aataaaatct gacatgtcaa araaaaaaaa mcy

## <u>12</u># 17

```
<211> 851
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 17..595
<221> sig_peptide
<222> 17..85
<223> Von Heijne matrix
      score 3.70000004768372
      seq FLPPLXRAFACRG/CQ
<221> polyA_signal
<222> 820..825
<221> polyA_site
<222> 840..851
<400> 243
                                                                       52
aagggggcgt ggggcc atg gtg gtc ttg cgg gcg ggg aag aag acc ttt ctc
                  Met Val Val Leu Arg Ala Gly Lys Lys Thr Phe Leu
                               -20
ccc cct ctm wgc cgc gcc ttc gcc tgc cgc ggc tgt caa ctc gct ccg
Pro Pro Leu Xaa Arg Ala Phe Ala Cys Arg Gly Cys Gln Leu Ala Pro
    -10
                        - 5
gag cgc ggc gcc gag cgc agg gat aca gcg ccc agc ggg gtc tca aga
                                                                       148
Glu Arg Gly Ala Glu Arg Arg Asp Thr Ala Pro Ser Gly Val Ser Arg
                                     15
                10
                                                                       196
ttc tgc cct cca aga aag tct tgc cat gat tgg ata gga ccc cca gat
Phe Cys Pro Pro Arg Lys Ser Cys His Asp Trp Ile Gly Pro Pro Asp
                                 30
aaa tat tca aac ctt cga cct gtt cac ttt tac ata cct gaa aat gaa
                                                                       244
Lys Tyr Ser Asn Leu Arg Pro Val His Phe Tyr Ile Pro Glu Asn Glu
                             45
tot coa ttg gaa caa aag ott aga aaa tta aga caa gaa aca caa gaa
                                                                       292
Ser Pro Leu Glu Gln Lys Leu Arg Lys Leu Arg Gln Glu Thr Gln Glu
                         60
tgg aat caa cag ttc tgg gca aac cag aat ttg act ttt agt aag gaa
Trp Asn Gln Gln Phe Trp Ala Asn Gln Asn Leu Thr Phe Ser Lys Glu
                    75
                                                                       388
aaa gaa gaa ttt att cac tca aga cta aaa act aaa ggc ctg ggc ctg
Lys Glu Glu Phe Ile His Ser Arg Leu Lys Thr Lys Gly Leu Gly Leu
                90
                                     95
aga act gaa tca ggt cag aaa gca aca ttg aat gca gaa gaa atg gcg
                                                                       436
Arg Thr Glu Ser Gly Gln Lys Ala Thr Leu Asn Ala Glu Glu Met Ala
                                 110
gac ttc tac aag gaa ttt tta agt aaa aat ttt cag aag cac atg tat
                                                                       484
Asp Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Gln Lys His Met Tyr
                                                 130
                             125
                                                                       532
tat aac aga gat tgg tac aag cgc aat ttt gcc atc acc ttc ttc atg
Tyr Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile Thr Phe Phe Met
                         140
                                                                       580
gga aaa gtg gcc ctg gaa agg att tgg aac aag ctt aaa cag aaa caa
 Gly Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu Lys Gln Lys Gln
                                         160
                     155
                                                                       635
 aag aag agg agc aac taggagtcca ctctgaccca gccagagtcc aggtttccac
 Lys Lys Arg Ser Asn
                 170
                                                                       695
 aggaagcara tggagctcct ttcacagggg ctctgagaaa aactggagct gatctcaaga
 agccccacat cttcctaagg ggccccatgg cctgtttggg ggcagggtag gtcctggggc
```

<221> sig_peptide

actgtgggcc gcctgcctgc tgatgtgggc tctaggccag cttgttgtca cgtacgtggt gtgaaataaa gcccaagcac tgggaaaaaa aaaaaa	815 851
<210> 244 <211> 495 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 89334	
<221> sig_peptide <222> 89130 <223> Von Heijne matrix score 3.59999990463257 seq AFTLXSLLQAALL/CV	
<221> polyA_signal	
<221> polyA_site <222> 484495	
<pre>&lt;400&gt; 244 agtaggaasg cgccgsccgt ggaggcgcca cgtcccttgc sgcggcggga gagamatcgc ttggacttcg gggcggcctc ggacggcc atg gcc ttt acc ctg tas tca ctg</pre>	60 112
ctg cag gca gcc ctg ctc tgc gtc aac gcc atc gca gtg ctg cac gag Leu Gln Ala Ala Leu Leu Cys Val Asn Ala Ile Ala Val Leu His Glu -5 10	160
gag cga ttc ctc aag aac att ggc tgg gga aca gac cag gga att ggt Glu Arg Phe Leu Lys Asn Ile Gly Trp Gly Thr Asp Gln Gly Ile Gly 15 20 25	208
gga ttt gga gaa gag ccg gga att aaa tca sag sta atg avs ctt att Gly Phe Gly Glu Glu Pro Gly Ile Lys Ser Xaa Xaa Met Xaa Leu Ile 30 35 40	256
cga tct gta aga acc gtg atg aga gtg cca ttg ata ata gta aac tca Arg Ser Val Arg Thr Val Met Arg Val Pro Leu Ile Ile Val Asn Ser 45 50 55	304
att gca att gtg tta ctt tta tta ttt gga tgaatwtcat tggagaaaat Ile Ala Ile Val Leu Leu Leu Phe Gly 60 65	354
ggakactcag aaraggacat gccaktaraa kttattactt tggtcattat tggaatattt atatcttagc tggctgacct tgcacttgtc aaaaatgtaa agctgaaaat aaaaccaggg tttctattta aaaaaaaaa a	414 474 495
<210> 245 <211> 884 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 21614	

<222> 21..83 <223> Von Heijne matrix score 10 seq LWALAMVTRPASA/AP

<221> polyA_signal <222> 849..854

<221> polyA_site <222> 873..884

<400> 245

<400> 245			
aataccttag accctcagtc atg cca gtg cct gct  Met Pro Val Pro Ala  -20	ctg tgc ctg ctc tgg gcc 53 Leu Cys Leu Leu Trp Ala -15		
ctg gca atg gtg acc cgg cct gcc tca gcg g Leu Ala Met Val Thr Arg Pro Ala Ser Ala A	cc ccc atg ggc ggc cca 101 la Pro Met Gly Gly Pro		
gaa ctg gca cag cat gag gag ctg acc ctg c Glu Leu Ala Gln His Glu Glu Leu Thr Leu L 10 15			
cag ctg ggc cag gcc ctc aac ggt gtg tac a Gln Leu Gly Gln Ala Leu Asn Gly Val Tyr A 25 30			
ctg aca aag gcc agg aac agc ctg ggt ctc t Leu Thr Lys Ala Arg Asn Ser Leu Gly Leu T 40 45			
ctc ctg ggg cag gag gtc agc cgg ggc cgg g Leu Leu Gly Gln Glu Val Ser Arg Gly Arg A 55 60 6			
cgg gca agc ctg ttg gaa act car atg gag g Arg Ala Ser Leu Leu Glu Thr Gln Met Glu G 75 80			
cag gca rag gcc aca gct gag gtg ctg ggg g Gln Ala Xaa Ala Thr Ala Glu Val Leu Gly G 90 95	lu Val Ala Gln Ala Gln 100		
aag gtg cta cgg gac agc gtg cag cgg cta d Lys Val Leu Arg Asp Ser Val Gln Arg Leu X 105			
gcc tgg ctg ggc cct gcc tac cga aaa ttt g Ala Trp Leu Gly Pro Ala Tyr Arg Lys Phe G 120	lu Val Leu Lys Ala Pro 130		
	eu Thr Gly His Val Xaa 45 150		
cgg car arg cgg gar atg gtg gca cag cag c Arg Gln Xaa Arg Glu Met Val Ala Gln Gln X 155	aa Xaa Leu Xaa Gln Ile 165		
cag gar aaa ctc cac aca gcg gcg ctc cca g Gln Glu Lys Leu His Thr Ala Ala Leu Pro A 170 175	la		
tgaggaccaa tcatgctgca aggaacactt ccacgccc			
gagetgeetg tteactggga teagecaggg egeeggge	• • •		
agacagacgc aggcggggac aaaggcagag gatgtagc	<del></del>		
aggacatgta ccctttcatr mctacacacc cctcatta	aa gcavagtegt ggcateteaa 874 884		
aaaaaaaaa	884		

<210> 246

<211> 897

<212> DNA

<213> Homo sapiens

<220> <221> CDS <222> 94..573 <221> sig_peptide <222> 94..258 <223> Von Heijne matrix score 4.69999980926514 seq IGILCSLLGTVLL/WV <221> polyA_signal <222> 862..867 <221> polyA_site <222> 886..897 <400> 246 60 aagggcggct gcctagcacc cggaagagcc gtcaacttag cgagcgcaac aggctgccgc 114 tgaggagctg gagctggtgg ggactgggcc gca atg gac aag ctg aag aag gtg Met Asp Lys Leu Lys Lys Val -55 ctg age ggg cag gac acg gag gac egg age etg tee gag gtt gtt 162 Leu Ser Gly Gln Asp Thr Glu Asp Arg Ser Gly Leu Ser Glu Val Val -45 -40 gag gca tot toa tta ago tgg agt acc agg ata aaa ggc tto att gcg 210 Glu Ala Ser Ser Leu Ser Trp Ser Thr Arg Ile Lys Gly Phe Ile Ala -25 -20 -30 tgt ttt gct ata gga att ctc tgc tca ctg ctg ggt act gtt ctg ctg 258 Cys Phe Ala Ile Gly Ile Leu Cys Ser Leu Leu Gly Thr Val Leu Leu -10 306 tgg gtg ccc agg aag gga cta cac ctc ttc gca gtg ttt tat acc ttt Trp Val Pro Arg Lys Gly Leu His Leu Phe Ala Val Phe Tyr Thr Phe 10 ggt aat atc gca tca att ggg agt acc atc ttc ctc atg gga cca gtg 354 Gly Asn Ile Ala Ser Ile Gly Ser Thr Ile Phe Leu Met Gly Pro Val 30 25 402 aaa cag ctg aag cga atg ttt gag cct act cgt ttg att gca act atc Lys Gln Leu Lys Arg Met Phe Glu Pro Thr Arg Leu Ile Ala Thr Ile 40 450 atg gtg ctg ttg tgt ttt gca ctt acc ctg tgt tct gcc ttt tgg tgg Met Val Leu Cys Phe Ala Leu Thr Leu Cys Ser Ala Phe Trp Trp 55 60 cat aac aag gga ctt gca ctt atc ttc tgc att ttg cag tct ttg gca 498 His Asn Lys Gly Leu Ala Leu Ile Phe Cys Ile Leu Gln Ser Leu Ala 70 ttg acg tgg tac agc ctt tcc ttc ata cca ttt gca agg gat gct gtg 546 Leu Thr Trp Tyr Ser Leu Ser Phe Ile Pro Phe Ala Arg Asp Ala Val 95 85 593 aaa aad tgt ttt gcc gtg tgt ctt gca taattcatgg ccagttttat Lys Xaa Cys Phe Ala Val Cys Leu Ala gaagetttgg aaggeactat ggacagaage tggtggacag ttttgtwact atettegaaa 653 713 cctctgtctt acagacatgt gccttttatc ttgcagcaat gtgttgcttg tgattcgaac 773 atttgagggt tacttttgga agcaacaata cattctcgaa cctgaatgtc agtagcacag gatgagaagt gggttctgta tcttgtggag tggaatcttc ctcatgtacc tgtttcctct 833 ctggatgttg tcccactgaa ttcccatgaa tacaaaccta ttcagcaaca gcaaaaaaaa 893 897 aaaa

157

205

253

301

349

397

457

517 518

```
WO 99/31236
<210> 247
<211> 518
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 74..397
<221> sig_peptide
<222> 74..127
<223> Von Heijne matrix
     score 7.69999980926514
      seq LLLLPVLGLLVSS/KT
<221> polyA_signal
<222> 472..477
<221> polyA_site
<222> 507..518
<400> 247
aaagaaagag ctgcsgtgca ggaattcgtg tgccggattt ggttagctga gcccaccgag
aggegeetge agg atg aaa get ete tgt ete ete ete eet gte etg
               Met Lys Ala Leu Cys Leu Leu Leu Pro Val Leu
                           -15
ggg ctg ttg gtg tct agc aag acc ctg tgc tcc atg gaa gaa gcc atc
Gly Leu Leu Val Ser Ser Lys Thr Leu Cys Ser Met Glu Glu Ala Ile
                        1
aat gag agg atc cag gag gtc gcc ggc tcc cta ata ttt agg gca ata
Asn Glu Arg Ile Gln Glu Val Ala Gly Ser Leu Ile Phe Arg Ala Ile
                15
age age att gge ega ggg age gag age gte ace tee agg ggg gae etg
Ser Ser Ile Gly Arg Gly Ser Glu Ser Val Thr Ser Arg Gly Asp Leu
                                35
get act tgc ccc cga ggc ttc gcc gtc acc ggc tgc act tgt ggc tcc
```

Ala Thr Cys Pro Arg Gly Phe Ala Val Thr Gly Cys Thr Cys Gly Ser 50

gcc tgt ggc tcg tgg gat gtg cgc gcc gag acc aca tgt cac tgc cag Ala Cys Gly Ser Trp Asp Val Arg Ala Glu Thr Thr Cys His Cys Gln

tgc gcg ggc atg gac tgg acc gga gcg cgc tgc tgt cgt gtg cag ccc Cys Ala Gly Met Asp Trp Thr Gly Ala Arg Cys Cys Arg Val Gln Pro

tgaggtcgcg cgcagcgcgt gcacagcgcg ggcggaggcg gctccaggtc cggaggggtt

65

80

<210> 248 <211> 350 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 51..242 <221> sig_peptide <222> 51..116 <223> Von Heijne matrix score 6.5
seq SCLCPALFPGTSS/FI

<221> polyA_signal <222> 319..324

<221> polyA_site <222> 339..350

<400> 248 56 acgtcattcc aaaaccacac ccttgcaaag ctttgtactc cgcaccccag atg atc tcc agg cag ctc aga tct ctt tcc tgc ctt tgc cct gca ctg ttc ccc 104 Ser Arg Gln Leu Arg Ser Leu Ser Cys Leu Cys Pro Ala Leu Phe Pro -10 -15 ggt act tcc tcc ttt att gta gca ctc agc tcc cca gcc gat ctg tac 152 Gly Thr Ser Ser Phe Ile Val Ala Leu Ser Ser Pro Ala Asp Leu Tyr 200 atc cct cav agg cas cga tct gat gaa ttg gtt ttt gaa tcc car aaa Ile Pro Xaa Arg Xaa Arg Ser Asp Glu Leu Val Phe Glu Ser Gln Lys 20 15 242 ggg tot gcc atg gag ttg gca gtc atc acg gta rat ggc gta Gly Ser Ala Met Glu Leu Ala Val Ile Thr Val Xaa Gly Val tgattttgct gaattttaaa taaaatgaaa accataaatt acatratgct tttattgach 302 350 cttgacmact ggcctaaata aaaaractct gactccaaaa aaaaaaaa

<210> 249

<211> 996

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 111..191

<221> sig_peptide

<222> 111..155

<223> Von Heijne matrix score 5.80000019073486 seq FLXLMTLTTHVHS/SA

<221> polyA_signal

<222> 965..970

<221> polyA_site

<222> 986..996

<400> 249

atccgataca gaacatgcag taatgtggac tgcccaccag aagcaggtga tttccgagct
cagcaatgct cagctcataa tgatgtcaag caccatggcc agttttatga atg ggy

Met Gly

-15

ttc ctg wgt cta atg acc ctg aca acc cat gtt cac tca agt gcc aag

Phe Leu Xaa Leu Met Thr Leu Thr Thr His Val His Ser Ser Ala Lys

-10

-5

1

cca aat gaa caa ccc tgg ttg ttg aac tagcacctaa ggtcttarat

Pro Asn Glu Gln Pro Trp Leu Leu Asn

5

10

271

ggtacgcgtt gctatacaga atctttggat atgtgcatca gtggtttatg ccaaattgtt

ggctgcgatc accagctggg aagcaccgtc aaggaarata actgtggggt ctgcaacrga

tc: ggi cti gai tai tc: gai tai	rgggt rgatg reasct casct rtegg rtegg ggctg recag raggt	ata gata gatc gaga gact gaga gaga gaga g	cctg act cag tac ccgg gcta acat tcaa	gccg tggt tata gaac tgag ctga attt acga tcaa tca	gct tgct tttt aag tct tct acg tct tcc	ggtc aatt ggaa cctt ggct taca tcct gagg caaa tagt	cgag ccct rcca gtgg ggac gtcc tgct agca ccca tcat	gg c gat a a c a c a c a c a c c a c c a c c a c c a c c a c c c c c c c c c c c c c c c c c c c c	arta gaag ccct attcac tcac kcat caac gtgt ttcac	taaa taka ccag tagt agcag cttc ctgt ggtt gggagt aaaca	t cc at c gg gg at at gg ct; gc at at a	cakc attc acta gact ttca caac gacc actc	tctc gcct awgg tccat tcgtta aagga tagga	cgc tgt tga gaa caa cat tca ctg	caacrga aaccaaa cttaaaa aaacagt wtttcca wattcgt ccaccga ccactat ctgtcca aaatagg caaaaaa		331 391 451 511 571 631 691 751 811 931 991
<21 <21	.0> 2 .1> 8 .2> D .3> H	60 NA	sapi	lens													
	0 > 1 > C		02														
<22	2> 4 3> V s	51 on H core	eijn 8.5	ie ma													
<22	2 > 8	28 olyA	_sit														
			860														
	0> 2! tata		acga	ggct	gc c	ggct	tagg	a cc	ccca	gctc	cga		t Se	r Pr	c tct o Ser		56
ggt Gly	Arg	Leu	Cys	Leu	Leu	acc Thr	Ile	Val	ggc Gly	ctg Leu	att Ile	ctc Leu	-2 ccc Pro	acc	aga Arg	1	104
gga Gly	cag	acg	ttg	aaa	gat	acc	acg	tcc	agt Ser	tct Ser 10	tca S <b>e</b> r	gca Ala	gac Asp	tca Ser	act Thr 15	]	L52
atc Ile	atg Met	gac Asp	att Ile	cag Gln 20	gtc Val	ccg Pro	aca Thr	cga Arg	gcc Ala 25	cca Pro	gat Asp	gca Ala	gtc Val	tac Tyr 30	aca	2	200
gaa Glu	ctc Leu	cag Gln	ccc Pro 35	acc Thr	tct Ser	cca Pro	acc Thr	cca Pro 40	acc	tgg Trp	cct Pro	gct Ala	gat Asp 45	gaa	aca Thr	2	48
cca Pro	caa Gln	ccc Pro 50	cag Gln	acc Thr	cag Gln	acc Thr	cag Gln 55	caa	ctg Leu	gaa Glu	gga Gly	acg Thr 60	gat Asp	gly aaa	cct Pro	2	96
cta Leu	gtg Val 65	aca Thr	gat Asp	cca Pro	gag Glu	aca Thr 70	cac	wak Xaa	agc Ser	mcc Xaa	aaa Lys 75	qca	gct Ala	cat His	ccc Pro	3	44
act Thr 80	gat Asp	gac Asp	acc Thr	acg Thr	acg Thr 85	ctc	tct Ser	gag Glu	aga Arg	cca Pro 90	tcc	cca Pro	agc Ser	aca Thr	kac Xaa 95	3	92

gtc cat dac aga ccb cba kda ccc tca akc cat ctg gtt ttc atg agg Val His Xaa Arg Pro Xaa Xaa Pro Ser Xaa His Leu Val Phe Met Arg 100 105	440
atg acc cct tct tct atg atg aac aca ccc tcc gga aac sgg ggc tgt Met Thr Pro Ser Ser Met Met Asn Thr Pro Ser Gly Asn Xaa Gly Cys 115 120 125	488
tgg tcg cag ctg tgc tgt tca tca cag gca tca tca tcc tca cca gtg Trp Ser Gln Leu Cys Cys Ser Ser Gln Ala Ser Ser Ser Pro Val 130 135 140	536
gca agt gca ggc agc tgt ccc ggt tat gcc gga atc att gca ggt gag Ala Ser Ala Gly Ser Cys Pro Gly Tyr Ala Gly Ile Ile Ala Gly Glu	584
tcc atc aga aac agg agc tgacaacctg ctgggcaccc gaagaccaag Ser Ile Arg Asn Arg Ser 160 165	632
ccccctgcca gctcaccgtg cccagcctcc tgcatcccct cgaagagcct ggccagagag ggaagacaca gatgatgaag ctggagccag ggctgccggt ccgagtctcc tacctccccc aaccctgccc gcccctgaag gctacctggc gccttggggg ctgtccctca agttatctcc tctgctaaga caaaaagtaa agcactgtgg tctttgcaaa aaaaaaaa	692 752 812 860
<210> 251 <211> 593 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 24560	
<221> sig_peptide <222> 24101 <223> Von Heijne matrix score 10.3999996185303 seq LLLLLCGPSQDQC/RP	·
<221> polyA_signal	
<221> polyA_site <222> 583593	
<400> 251	
aanccagctg csgccggcca gcc atg gag act gga gcg ctg cgg cgc ccg caa Met Glu Thr Gly Ala Leu Arg Arg Pro Gln -25 -20	53
ctt ctc ccg ttg ctg ctg ctc tgc ggc cct tcc cag gat caa tgc Leu Leu Pro Leu Leu Leu Leu Cys Gly Pro Ser Gln Asp Gln Cys -15 -10 -5	101
cga cct gta ctc cag aat ctg ttg cag agc cca ggc ttg aca tgg agc Arg Pro Val Leu Gln Asn Leu Leu Gln Ser Pro Gly Leu Thr Trp Ser 1 5 10 15	149
ttg gaa gtg ccc act ggg aga gaa gga aag gaa ggt ggg gat cgg gga Leu Glu Val Pro Thr Gly Arg Glu Gly Lys Glu Gly Gly Asp Arg Gly 20 25 30	197
CCa ggg cta akt ggg gcc act cca gcc agg agc cct cag ggc aag gag Pro Gly Leu Xaa Gly Ala Thr Pro Ala Arg Ser Pro Gln Gly Lys Glu 35 40 45	245
atg ggg aga caa agg acc aga aag gtg aag ggc cct gct tgg akt cac Met Gly Arg Gln Arg Thr Arg Lys Val Lys Gly Pro Ala Trp Xaa His 50 55 60	293

	•																
7	Chr	gca Ala	aat Asn	cag Gln	gaa Glu	cta Leu 70	aac Asn	agg Arg	atg Met	agg Arg	tct Ser 75	ctg Leu	tct Ser	tct Ser	GIA	tcc Ser 80	341
	55 gtg /al	cca Pro	gtg Val	Gly ggg	His	ctq	gag Glu	ggt Gly	ggc Gly	Thr	gtc	aag Lys	ctt Leu	cag Gln	aag	gac	389
a	acg Thr	ggc Gly	ctc Leu	His	85 tcc Ser	tgc Cys	ara Xaa	gat Asp	ggt Gly	90 atg Met	gct Ala	tct Ser	ctt Leu	gaa Glu 110	999	acg Thr	437
I	cca Pro	gct Ala	Ser	100 gtc Val	ctg Leu	gct Ala	gat Asp	Ala	105 tgc Cys	cca Pro	gga Gly	ttc Phe	cat His 125	gat	gtg Val	aan Xaa	485
9	gtt Val	Gln	115 arg Xaa	gcc Ala	cta Leu	ttt Phe	Gly	120 tta Leu	agt Ser	ggg Gly	ana Xaa	rta Xaa 140	ctg	tgg Trp	ctg Leu	aaa Lys	533
•	Thr	130 cac His	ttc Phe	tgc Cys	ctt Leu	Ser	135 att Ile	ana Xaa	ctt Leu	taaa	ataaa		ctgaa	arac	ct		580
	145 gtaa	aaaa	aaa a	aaa		150											593
	<210	)> 2	52														
	<212	l> 1 2> D 3> H		sapi	ens												
	<220	O >															
		l> C 2> 1	DS 09!	558								•					
	<22	2> 1 3> V s	core	273 eijn 3.7	de e ma 0000 TLPI	0047	6837	2									
		-	olyA 104.	_													
	att	0> 2 agct	stc	caag	gtct	cc c	ccag	cact	g ag	gagc	tcgc	ctg	ctgc	cct	cttg	cgcgcg	60
	gga	agca	gca	ccaa	gttc	ac g	gcca	acgc	c tt	ggca	ctag	ggt	ccag	a at	g gc t Al	t aca a Thr	117
	aca Thr	gto Val	cct Pro	Asp	ggt Gly	tgc Cys	cgc	aat Asn -45	Gly	ctg Leu	aaa Lys	tcc Ser	aag Lys -40	tac Tyr	tac	aga Arg	165
	ctt Leu	tgt Cys	gat Asp	aad	gct Ala	gaa Glu	gct Ala -30	Trp	ggc Gly	atc Ile	gtc Val	cta Leu -25	Glu	acg Thr	gtg Val	gcc Ala	213
	aca Thr	gcc	: aaa	gtt Val	gtg Val	acc Thr	Ser	gtg Val	gcc Ala	ttc Phe	atg Met -10	Leu	act Thr	cto Lev	ccg Pro	atc Ile -5	261
	ctc	ato	tgo Cys	aag Lys	gtg Val	cac	gac	tcc Ser	aac Asn	agg Arg	cga	aaa	atg Met	cto Lev	cct Pro	act Thr	309
	cag Gln	ttt Phe	e Leu	ttc Phe	ctc Leu	ctg Lev	ggt Gly	v Val	ttg Leu	ggc Gly	ato Ile	ttt Phe	: Gly	cto	acc Thr	ttc Phe	357
	gcc Ala	tto Phe	15 ato Ile	ato	gga Gly	cto Lev	gac Asp	20 : ggs : Gly	g ago / Ser	aca Thr	ggg	ccc Pro	25 aca Thr	cgc Arg	tto Phe	ttc Phe	405

25	
30 35 40 ctc ttt ggg atc ctc ttt tcc atc tgc ttc tcc tgc ctg ctg gct cat.	453
Leu Phe Gly Ile Leu Phe Ser Ile Cys Phe Ser Cys Leu Leu Ala His	•
45 50 55 60	E01
gct gtc agt ctg acc aag ctc gtc cgg ggg agg aaa gcc cct ttc cct Ala Val Ser Leu Thr Lys Leu Val Arg Gly Arg Lys Ala Pro Phe Pro	501
65 70 75	
gtt ggt gat tot ggg tot ggc ogt ggg ott oag oot agt ooa gga tgt	549
Val Gly Asp Ser Gly Ser Gly Arg Gly Leu Gln Pro Ser Pro Gly Cys	
80 85 90	598
tat cgc tat tgaatatatt gtcctgacca tgaataggac caacgtcaat Tyr Arg Tyr	570
95	
gtottttotg agotttocgo tootogtogo aatgaaaact ttgtoctoot gotoacotac	658
kteetettet tgatggeget gacetteete wtgteeteet teacettetg tggtkeette	718 778
acgggctgga avagacatgg ggcccacatc tacctcasga tgctcskctc cattgccatc tgggtggcct ggatcaccct gctcatgctt cctgactttg accgcrggtg ggatgacacc	838
atemtearet eegeettggs tresaatgge tgggtgttee tgttggetta tgttagteee	898
gagttttggc tgctcacaaa gcaackaaac cccatggatt atcctgttga ggatgctttc	958
tgtaaacctc aactcgtgaa gaagagctat ggtgtggrga acagagccta skctcaagag	1018
gaaatcactc aaggttttga agagacaggg gacacgctct atgcccccta ttccacacat	1078
tttcagctgc agaascagcc tccccaaaaa aaaaaa	1114
<210> 253	
<211> 1182	
<212> DNA	
<213> Homo sapiens	,
<220>	
<221> CDS	
<222> 128835	
<221> sig_peptide	
<222> 128220	
<223> Von Heijne matrix	
score 4.69999980926514	
seq LAVDSWWLDPGHA/AV	
<221> polyA signal	
<222> 11451150	
<221> polyA_site	
<222> 11701181	
<400> 253	
aagaactgcg tctcgcgacc caggcgcggg ttcccggagg acagccaaca agcgatgctg	60
ccgccgccgt ttcctgattg gttgtgggtg gctacctctt cgttctgatt ggccgctagt	120
gagcaag atg ctg agc aag ggt ctg aag cgg aaa cgg gag gag gag gag	169
Met Leu Ser Lys Gly Leu Lys Arg Lys Arg Glu Glu Glu -30 -25 -20	
gag aag gaa cot otg goa gto gao too tgg tgg ota gat cot ggo cac	217
Glu Lys Glu Pro Leu Ala Val Asp Ser Trp Trp Leu Asp Pro Gly His	
-15 -10 -5	
gea geg gtg gea cag gea eee eeg gee gtg gee tet age tee ete ttt	265
Ala Ala Val Ala Gln Ala Pro Pro Ala Val Ala Ser Ser Ser Leu Phe 1 5 10 15	
gac ctc tea gtg ctc aag ctc cac cac agc ctg cag vrr agt rag ccg	313
Asp Leu Ser Val Leu Lys Leu His His Ser Leu Gln Xaa Ser Xaa Pro	
20 25 30	2.52
gac ctg cgg cac ctg gtg ctg gtc atr aac act ctg cgg cgc atc cag	361

_		_	35					40					45	Ile	-	
gcg Ala	tcc Ser	atg Met 50	gca Ala	ccc Pro	gcg Ala	gct Ala	gcc Ala 55	ctg Leu	cca Pro	cct Pro	gtg Val	cct Pro 60	acc Thr	cca Pro	cct Pro	409
gca Ala	gcc Ala 65	ccc Pro	ant Xaa	gtg Val	gct Ala	gac Asp 70	aac Asn	tta Leu	ctg Leu	gca Ala	agc Ser 75	tcg Ser	gac Asp	gct Ala	gcc Ala	457
ctt Leu 80	tca Ser	gcc Ala	tcc Ser	atg Met	gcc Ala 85	arm Xaa	ctc Leu	ctg Leu	gar Glu	gac Asp 90	ctc Leu	agc Ser	cac His	att Ile	gag Glu 95	505
ggc	ctg Leu	agt Ser	cag Gln	gct Ala 100	ccc Pro	caa Gln	ccc Pro	ttg Leu	gca Ala 105	gac Asp	gag Glu	Gly 999	cca Pro	cca Pro 110	ggc Gly	553
cgt Arg	agc Ser	Ile	999 Gly	gga	wca Xaa	ccg Pro	ccc Pro	amc Xaa 120	ctg Leu	ggt Gly	gcc Ala	ttg Leu	gac Asp 125	ctg Leu	ctg Leu	601
ggc Gly	cca Pro	gcc	act	ggc Gly	tgt Cys	cta Leu	ctg Leu 135	gac Asp	aat Asn	gly ggg	ctt Leu	gag Glu 140	ggc	ctg Leu	ttt Phe	649
gag Glu	gat Asp 145	att	gac Asp	acc Thr	tct Ser	atg Met 150	tat	gac Asp	aat Asn	gaa Glu	ctt Leu 155	tgg Trp	gca Ala	cca Pro	gcc Ala	697
tct Ser 160	gag	ggc Gly	ctc Leu	aaa Lys	cca Pro 165	ggc	cct Pro	gag Glu	gat Asp	999 Gly 170	ccg	Gly	aag Lys	gag Glu	gaa Glu 175	745
gct	ccg Pro	gag Glu	ctg Leu	gac Asp 180	gag	gcc Ala	gaa Glu	ttg Leu	gac Asp 185	tac	ctc Leu	atg Met	gat <b>Asp</b>	gtg Val 190	ctg Leu	793
gtg Val	ggc Gly	aca Thr	cag Gln 195	gca	ctg Leu	gag Glu	cga Arg	ccg Pro 200	ccg	G] À 333	cca Pro	Gly 999	cgc Arg 205			835
tgt gag ggg ctg	cctc agac gagc gccc	gaa aga ctg tgg	tgct aaga atct gaat ggtc	cacag agtc tacc atag	gc to ct go cc co ct to	ggcti ggca tagti gggci	tccci actto gatgo	t ato	taca atcc tgac tctc	gaga gtcc aggg tgat	tcc; tct; acg;	ggaa ggaa ggaa	tgg ( ttg ( tca (	ggcc: gggc! actg:	accaac actttg tggcag aattcc aatcag	895 955 1015 1075 1135 1182

<211> 1073

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 59..505

<221> sig_peptide

<222> 59..358

<223> Von Heijne matrix score 3.70000004768372 seq LASSFLFTMGGLG/FI

<221> polyA_signal

<222> 1042..1047

```
<400> 254
actgtttnng ggaggcgcgt ggggcttgag gccgagaacg gcccttgctg ccaccaac.
                                                                      58
atg gag act ttg tac cgt gtc ccg ttc tta gtg ctc gaa tgt ccc aac
                                                                     106
Met Glu Thr Leu Tyr Arg Val Pro Phe Leu Val Leu Glu Cys Pro Asn
                                        -90
                    -95
ctg aag ctg aag aag ccg ccc tgg ttg cac atg ccg tcg gcc atg act
Leu Lys Leu Lys Lys Pro Pro Trp Leu His Met Pro Ser Ala Met Thr
                -80
                                    -75
                                                                     202
gtg tat gct ctg gtg gtg tct tac ttc ctc atc acc gga gga ata
Val Tyr Ala Leu Val Val Val Ser Tyr Phe Leu Ile Thr Gly Gly Ile
                                -60
att tat gat gtt att gtt gaa cct cca agt gtc ggt tct atg act gat
                                                                     250
Ile Tyr Asp Val Ile Val Glu Pro Pro Ser Val Gly Ser Met Thr Asp
                            -45
gaa cat ggg cat cag agg cca gta gct ttc ttg gcc tac aga gta aat
                                                                     298
Glu His Gly His Gln Arg Pro Val Ala Phe Leu Ala Tyr Arg Val Asn
                      -30
gga caa tat att atg gaa gga ctt gca tcc agc ttc cta ttt aca atg
                                                                     346
Gly Gln Tyr Ile Met Glu Gly Leu Ala Ser Ser Phe Leu Phe Thr Met
-20
                   -15
                                        -10
                                                                     394
gga ggt tta ggt ttc ata atc ctg gac gga tcg aat gca cca aat atc
Gly Gly Leu Gly Phe Ile Ile Leu Asp Gly Ser Asn Ala Pro Asn Ile
                                                                     442
cca aaa ctc aat aga ttc ctt ctt ctg ttc att gga ttc gtc tgt gtc:
Pro Lys Leu Asn Arg Phe Leu Leu Phe Ile Gly Phe Val Cys Val
       15
                            20
                                                                     490
cta twr agt ttt tkc ayg gct aga gta ttc atg aga atg aaa ctg ccg
Leu Xaa Ser Phe Xaa Xaa Ala Arg Val Phe Met Arg Met Lys Leu Pro
ggc tat ctg atg ggt tagagtgcct ttgasaagaa atcagtggat actggatttg
                                                                     545
Gly Tyr Leu Met Gly
                                                                     605
ctcctgtcaa wgaastttta aaggctgtmc caatcctcta atatgaaatg tggaaaagaa
                                                                     665
tgaagagcag cagtaaaaga aatatctagt gaaaaaacag gaagcgtatt gaagcttgga
ctagaatttc ttcttggtat taaagagaca agtttatcac agaatttttt ttcctgctgg
                                                                     725
cctattgcta taccaatgat gttgagtggc attttctttt tagtttttca ttaaaatata
                                                                     785
ttccatatct acaactataa tatcaaataa agtgattatt ttttacaacc ctcttaacat
                                                                     845
tttttggaga tgacatttct gattttcaga aattaacata aaatccagaa gcaagattcc
                                                                     905
                                                                     965
gtaagctgag aactctggac agttgatcag ctttacctat ggtgctttgc ctttaactag
agtgtgtgat ggtagattat ttcagatatg tatgtaaaac tgtttcctga acaataagat
                                                                    1025
gtatgaacgg agcagaaata aatacttttt ctaattaaaa aaaaaaaa
                                                                    1073
```

<211> 818

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 1..207

<221> sig peptide

<222> 1..147

<223> Von Heijne matrix score 7.59999990463257 seg HLPFLLLLSCVGX/XP

<221> polyA_signal

<222> 784..789

<221> polyA_site <222> 807..818 <400> 255 atg cct ttc cat ttt ccg ttc ctt ggg ttt gtg tgt ctg cat ctc cat 48 Met Pro Phe His Phe Pro Phe Leu Gly Phe Val Cys Leu His Leu His -45 -40 ctt acc cct tgc ctg act gta ccc cgt aga ccc ctg ttt ctc ctc ctg 96 Leu Thr Pro Cys Leu Thr Val Pro Arg Arg Pro Leu Phe Leu Leu Leu -20 -30 -25 144 cac ctg tgt ccc cat ctg ccc ttc ttg ttg ctc ctg tca tgt gtc ggg His Leu Cys Pro His Leu Pro Phe Leu Leu Leu Ser Cys Val Gly -10 -15 192 gke www eec tee tgt etg eet tet tee tee act tgt gte age ttg eat Xaa Xaa Pro Ser Cys Leu Pro Ser Ser Ser Thr Cys Val Ser Leu His 10 ttt ttt att cct gac tgagtcacca cacccctctc ccctgatcaa agggaatatk 247 Phe Phe Ile Pro Asp 307 artttttaat ttggatcgac tgaggtgcca ggagaaactg cagkcccagg tatccmvaca gccaccagga tggtccctcg ccccaccccc accgcctctk ccccaccttt tccaacgtgt 367 tgcatgctgg gaactggggg gtgtggggga aggggctgcc ggcttctttc aggangctga 427 487 rgtttggarg caaaatcaac ctgggaracc accccggccg cggcgcctca gtggacaggt 547 gggargaaaa gaaaacttct taccttggar garggacatc ccgcttcctt atccttagct 607 tttttgttgc tcctccccac tgcccctttt aatttatttg gttgtttgcg gaaggagggg ggaaggggt aagctgggcc gggaactgtc cgaggtgctg agctggggcg ggaccggaat 667 727 cctcccggta gggtaccagg gactgagttg ggcctggggc cgtgtccaag gtgccaatga 787 tgcgggccga cagarcgggc cgcactgtct gtctgtccgt ctgtcccgga aagaactata aagcgctgga agcgcctgca aaaaaaaaaa a <210> 256 <211> 971 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 12..734 <221> sig_peptide <222> 12..101 <223> Von Heijne matrix score 4.80000019073486 seq ILFCVGAVGACTL/SV <221> polyA signal <222> 914..919 <221> polyA site <222> 961..971 <400> 256 aatacacaga a atg ggg act gcg agc aga agc aac atc gct cgc cat ctg 50 Met Gly Thr Ala Ser Arg Ser Asn Ile Ala Arg His Leu -25 -20 -30 caa acc aat ctc att cta ttt tgt gtc ggt gct gtg ggc gcc tgt act 98 Gln Thr Asn Leu Ile Leu Phe Cys Val Gly Ala Val Gly Ala Cys Thr - 5 ......-10 146 ctc tct gtc aca caa ccg tgg tac cta gaa gtg gac tac act cat gag

Leu Ser Val Thr Gln Pro Trp Tyr Leu Glu Val Asp Tyr Thr His Glu

	1				5					10					15	
acc	qtc	acc	ata	aag	tgt	acc	ttc	tcc	gca	acc	gga	tgc	cct	tct	gag	194
Ala	Val	Thr	Ile	Lys	Cys	Thr	Phe	Ser	Ala	Thr	Gly	Cys	Pro	Ser	Glu	
				20					25					30		
caa	cca	aca	tgc	ctg	tgg	ttt	cgc	tac	ggt	gct	cac	cag	cct	gag	aac	242
Gln	Pro	Thr	Cys	Leu	Trp	Phe	Arg	Tyr	Gly	Ala	His	Gln	Pro	Glu	Asn	
			35					40					45			
ctg	tgc	ttg	gac	999	tgc	aaa	agt	gag	gca	gas	aag	ttc	aca	gtg	agg	290
Leu	Cys	Leu	Asp	Gly	Cys	Lys	Ser	Glu	Ala	Xaa	Lys	Phe	Thr	Val	Arg	
		50					55					60				
gag	gcc	ctc	aaa	gaa	aac	caa	gtt	tcc	.ctc	act	gta	aac	aga	gtg	act	338
Glu	Ala	Leu	Lys	Glu	Asn	Gln	Val	Ser	Leu	Thr	Val	Asn	Arg	Val	Thr	
	65					70					75					206
tca	aat	gac	agt	gca	att	tac	atc	tgt	gga	ata	gca	ttc	CCC	agt	gtg	386
Ser	Asn	Asp	Ser	Ala	Ile	Tyr	Ile	Cys	Gly	Ile	Ala	Phe	Pro	Ser	vai	
80					85					90					95	424
ccg	gaa	gcg	aga	gct	aaa	cag	aca	gga	gga	aaa	acc	aca	ctg	gtg	gta	434
Pro	Glu	Ala	Arg	Ala	Lys	Gln	Thr	Gly		GIY	Thr	Thr	Leu	Val	vai	
				100					105					110		4.00
aga	gaa	att	aag	ctg	ctc	agc	aag	gaa	ctg	cgg	agc	ttc	ctg	aca	gct	482
Arg	Glu	Ile	Lys	Leu	Leu	Ser	Lys		Leu	Arg	ser	Phe	Leu	Thr	Ala	
			115					120					125			520
ctt	gta	tca	ctg	ctc	tct	gtc	tat	gtg	acc	ggt	gtg	tgc	gtg	gcc	DD-	530
Leu	Val	Ser	Leu	Leu	ser	Val		Val	Thr	Gly	Val	Cys	Val	Ala	Pne	
		130					135					140		~	2 + 2	578
ata	ctc	ctc	tcc	aaa	tca	aaa	tcc	aac	CCE	cta	aga	aac	dad	gaa	Tla	٥ / ر
Ile		Leu	Ser	Lys	Ser		ser	Asn	Pro	ьeu	arg	ASII	пуs	Glu	116	
	145					150					155	~ + +	+++	cad	C a a	626
aaa	gaa	gac	tca	caa	aag	aag	aag	agt Co~	get Nla	299	cgc	Tla	Dhe	cag	Glu	020
_	GIU	Asp	ser	GIN		ьys	гуя	ser	Ala	170	AT 9	110	F11C	Gln	175	
160					165			202	ant.		~~~	202	aat	cag		674
att	gct	caa	gaa	cta	tac	Cat	aay	aya	Uic	7723	Glu	Thr	Asn	cag Gln	Gln	•
ııe	Ala	GIN	GIU	180	IYL	urs	пуэ	nr 9	185	vai	Olu			190		
			~		220	3.C.t	+=+	as a		ara	aga	ata	ctt	tac	aac	722
Cot	gag	Tura	yat	Aac	Aac Acn	Thr	Tyr	Glu	Agn	Ara	Ara	Val	Leu	Ser	Asn	
ser	GIU	пув	195	ASII	ASII	1111	TYL	200	AUII		••••		205			
+ = +	~ = =	= ~~		tar	2220	<b>3</b> ++	ttaa		саа	tgaa	atca	c ta		tcca		774
	Glu				aaac	900	ccuu			0500	5					
TYT	Giu	210	110													
act	ccad		ctat	aaca	at a	ttaa	tgaa	c at	atat	catc	agq	tctt	aaa	aaaa	aataaa	834
aat	2220	taa	aaad	35 <b>0</b> 0	et a	acta	caaa	a aa	ggat	gcca	raa	tqta	agg	aaac	tataac	894
taa	takt:	cat	tacc	2222	ta c	taaa	accc	a ac	aaaa	tqca	act	gaaa	aat	acct	tccaaa	954
	gcca									J		_				971
	gcca	uaa	uuua	~~ w												

<211> 640

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 378..518

<221> sig_peptide

<222> 378..467

<223> Von Heijne matrix score 5.5 seq SLMTCTTLINASA/IS

<221> polyA_signal <222> 607..612 <221> polyA site 1 <222> 628..640 <400> 257 agcctgggta akgcccaaga tggctgtctt cgccttagta ctcgtgtgaa gttggcgggg 60 acggttcctg tcatcttctt gggcttattt ggtgtgctgt tgaagggggg agactagaga 120 aatggcaggg aacctcttat ccggggcagg taggcgcctg tgggactggg tgcctctggc 180 gtgcagaagc ttctctcttg gtgtgcctag attgatcggt ataaggctca ctctcccgcc 240 ccccaaagtg gttgatcgtt ggaacgagaa aagggccatg ttcggagtgt atgacaacat 300 cgggatcctg ggaaactttg aaaagcaccc caaagaactg atcagggggc ccatatggct 360 tcgaggttgg aaaggga atg aat tgc aac gtt gta tcc gaa aga gga aaa 410 Met Asn Cys Asn Val Val Ser Glu Arg Gly Lys -30 -25 tgg ttg gaa gta gaa tgt tcg ctg atg acc tgc aca acc tta ata aac 458 Trp Leu Glu Val Glu Cys Ser Leu Met Thr Cys Thr Thr Leu Ile Asn -15 -10 gca tcc gct atc tct aca aac act tta acc gac atg gga agt ttc gat 506 Ala Ser Ala Ile Ser Thr Asn Thr Leu Thr Asp Met Gly Ser Phe Asp aga aga gaa agc tgagaacttc ggaaaaggct catctgtcac cctggaraag 558 Arg Arg Glu Ser ggaaactgta cttttccctg tgaggaaacg gctttgtatt ttctctgtaa taaaatgggg 618 cttctttgga aaaaaaaaa aa 640 <210> 258 <211> 745 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 110..304 <221> sig_peptide <222> 110..193 <223> Von Heijne matrix score 4.59999990463257 seq PLQWSLLVAVVAG/SV <221> polyA_signal <222> 708..713 <221> polyA_site <222> 732..743 <400> 258 actteegeet gegeetgege ageveagete eshgageeet gecaaceatg gtgaacttgg 60 gtctgtcccg ggtggacgac gccgtggctg ccaagcaccc ggcaccggc atg gcc ttt 118 Met Ala Phe ggc ttg cag atg ttc att cag agg aag ttt cca tac cct ttg cag tgg 166 Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro Leu Gln Trp -25 -20 -15 -10 age etc eta gtg gee gtg gtt gea gge tet gtg gte age tae ggg gtg 214

Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser Tyr Gly Val

acg aga gtg gag tog gag aaa tgo aac aac oto tgg oto tto otg gag

Thr Arg Val Glu Ser Glu Lys Cys Asn Asn Leu Trp Leu Phe Leu Glu	
acc gga cag ctc ccc aaa gac agg agc aca gat cag ara agc Thr Gly Gln Leu Pro Lys Asp Arg Ser Thr Asp Gln Xaa Ser	304
taggagaget ccagcagggg cacagargat tgggggcagg argartetgg aacacakeet	364
tratocccc tgaccccagg ccgaccctcc ccacacccta gggtacccca gtcgtatcct	424
cratecacat atatagecag acctaacaaa emeetgeaga tagetgetge decaadetag	484 544
gacctgccca ggaggttgga gcagaaaggg ctctccctgg ggtggtgttt ctcctctagg gtattgggat gcatgttctg cactgccagc agagagggtg tgtctggggg ccaccaccta	604
tgggacacgg ggtcgaaggg gcctgtacac tctgtcattt cctttctagc ccctgcatct	664
ccaacaagtc caaggtgaca gctggtgcta ggggcgtggg gttaataaat ggcttatcct	724
tctctccaaa araaaaaaam C	745
.<210> 259	
<211> 637	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 201419	
<221> sig_peptide	
<222> 201272	
<223> Von Heijne matrix	
score 6.4000009536743	
seq LSYLPLWLGPIWP/CS	
<221> polyA_signal	
<222> 601606	
201. maluh mita	
<221> polyA_site <222> 627637	*
(222) (21,103)	
<400> 259	60
acaaaatata attgeetets ceeteteeca ttttetetet tgggageaat ggteacagte cetggtacet gaaaaggtae etaggtetag geeettette cettteectt ceteteecet	120
acccagaac titggetece titecettet cietetgta getecaggag geetgtgate	180
cagetecetg ectageatee atg ace tgt tgg atg tta cet cea ate agt tte	233
Met Thr Cys Trp Met Leu Pro Pro Ile Ser Phe	
-20 -15	281
ctg tcc tac ctg cct ctt tgg ctt gga cct ata tgg cca tgc tct ggc Leu Ser Tyr Leu Pro Leu Trp Leu Gly Pro Ile Trp Pro Cys Ser Gly	
-10 -5 1	
tot acc off ggg aag oof gat ood ggt gtg tgg ood ago ftg tto agg	329
Ser Thr Leu Gly Lys Pro Asp Pro Gly Val Trp Pro Ser Leu Phe Arg	
5 10 15 15 and and are stated as a second	377
ccc tgg gat gct gca tct cca ggc aac tat gca ctt tcc cgg gga rar	5,,
Pro Trp Asp Ala Ala Ser Pro Gly Asn Tyr Ala Leu Ser Arg Gly Xaa 20 25 30 35	
aac cak tat gav aak tgg ggg cag ggc aca cat tca tct ttg	419
Asn Xaa Tyr Xaa Xaa Trp Gly Gln Gly Thr His Ser Ser Leu	
40 45	470
targaaggte tggcetgggg terggtgaag gagggeecag gteagttetg gggteecagt	479 539
gacctgcttt gccattctcc tggtgccgct gctgctccct gtttctggag ctggatgttc cccacctggc agttgagctg cctgagccaa tgtgtctgtc tttggtaact gagtgaacca	599
taataaaggg gaacatttgg ccctgtgaaa aaaaaaaa	637

```
<210> 260
<211> 1315
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 123..302
<221> sig_peptide
<222> 123..176
<223> Von Heijne matrix
     score 4.30000019073486
     seg WTCLKSFPSPTSS/HA
<221> polyA signal
<222> 1279..1284
<221> polyA site
<222> 1301..1312
<400> 260
aagagcatcc tgcgccccgg cgcggggccc tgcggtagcc tcaggcccct cccctggacc
                                                                    60
cgccgcagag ccagtgcaga atacagaaac tgcagccatg accacgcacg tcaccctgga
                                                                    120
ag atg ccc tgt cca acg tgg acc tgc ttg aag agc ttc ccc tcc ccg
                                                                    167
   Met Pro Cys Pro Thr Trp Thr Cys Leu Lys Ser Phe Pro Ser Pro
              -15
acc agc agc cat gca tog agc oto cac ott cot coa toa tgt acc agg
                                                                    215
Thr Ser Ser His Ala Ser Ser Leu His Leu Pro Pro Ser Cys Thr Arg
                                                                    263
cta act ttg aca caa act ttg agg aca gga atg cat ttg tca cgg gca
Leu Thr Leu Thr Gln Thr Leu Arg Thr Gly Met His Leu Ser Arg Ala
                                           25
                                                                    312
ttg caa ggt aca ttg acc agg cta cag tcc act cca gca tgaatgarat
Leu Gln Gly Thr Leu Thr Arg Leu Gln Ser Thr Pro Ala
                    35
gctggaggaa ggacatgakt atgcggtcat gctgtacacc tggcgcagct gttcccgggc
                                                                    372
                                                                    432
cattccccag gtgaaatgca acragcagcc caaccgakta raratctatg araaracagt
araggtgctg gagccggagg tcaccaagct catgaagttc atgtattttc arcgcaaggc
                                                                    492
categagegg ttetgeaseg aggtgaageg getgtgeeat geegagegea ggaaggaett
tgtctctgag gcctacctcc tgacccttgg caagttcatc aacatgtttg ctgtcctgga
                                                                    612
tgagctaaag aacatgaast gcagcgtcaa raatgaccac tctgcctaca agagggcagc
                                                                    672
acagtteetg eggaagatgg cagateecca gtetateeag gagtegeaga acettteeat
                                                                    732
gttcctggcc aaccacaaca ggatcaccca gtgtctccac cagcaacttg aagtgatccc
                                                                    792
aggctatgag gagctgctgg ctgacattgt caacatctgt gtggattact acgagaacaa
                                                                    852
gatgtacctg actcccagtg agaaacatat gctcctcaag gtaaaactcc cctgaggccg
                                                                    912
cacccatgga gcctgggctt accctctcac cttcttctta ttaaaaatcc gttttaaaaa
                                                                    972
                                                                   1032
acaatgtttc ttttttctta aacattgata cagatcttac ggcacataat ggtttgtaac
                                                                   1092
ctgttccttt cctgtaatat aatataccgt agtcaccttt ccagatgtca ttaaggctat
ttctacaatg ttatgtgtaa tgactgccaa gtattctgtt gtattggaac attgtcatgt
                                                                   1152
aacatatccc ctgtggttgg atatttgcta aacttcattg aacacccttg tagcagtttt
                                                                   1212
tgtgcacatc tttttgtcaa ggcaaacttc ctagaagaga aattgctggc tcaaagggaa
                                                                   1272
                                                                   1315
```

<210> 261

<211> 1035

<212> DNA

<213> Homma sapiens

PCT/IB98/02122

	L> CI	os 86	73													
<222	2> 98 3> Vo	B3' on He	eptio 76 eijne 5.59 LLLR(	e mai	9904		7									·
	-		_site .1035		· · · · · · · · · · · · · · · · · · ·											
aatt		ygt g								ate	g gca	a ga	g tt	g ggd u Gly	tggcct c cta y Leu	60 115
														gct Ala		163
tca Ser	aag Lys -70	aga Arg	ggc Gly	ttg Leu	aga Arg	ctc Leu -65	aaa Lys	act Thr	gta Val	gat Asp	tcc Ser -60	tgc Cys	ttc Phe	caa Gln	gac Asp	211
														gaa Glu		259
														gtg Val -25		307
														cga Arg		355
ctg Leu	ttt Phe	gca Ala -5	caa Gln	gct Ala	gag Glu	aag Lys	tgg Trp 1	tat Tyr	ctt Leu	aag Lys	cta Leu 5	cag Gln	aca Thr	gac Asp	atc Ile	403
tct Ser 10	gaa Glu	ctt Leu	gaa Glu	aac Asn	cga Arg 15	gaa Glu	tta Leu	tta Leu	gaa Glu	caa Gln 20	ktt Xaa	gca Ala	gaa Glu	ttt Phe	gaa Glu 25	451
aaa Lys	gca Ala	rav Xaa	att Ile	aca Thr 30	tct Ser	tca Ser	aac Asn	aaa Lys	aag Lys 35	ccc Pro	atc Ile	tta Leu	dat Xaa	gtc Val 40	aca Thr	499
														ctc Leu		547
														att Ile		595
														gtt Val		643
			agg Arg							tagi	ttt	act 1	tgat	ggta	cc	693
															gatcac	753
															acattt	813
															ttatc	873 933
															ggcagg gcactc	933
			gacag												,	1035

```
<210> 262
<211> 696
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 17..463
<221> sig peptide
<222> 17..232
<223> Von Heijne matrix
     score 3.79999995231628
     seq LMGLALAVYKCQS/MG
<221> polyA_signal
<222> 657..662
<221> polyA_site
<222> 684..696
<400> 262
                                                                      52
actcaaacag attccc atg aat ctc ttc atc atg tac atg gca ggc aat act
                  Met Asn Leu Phe Ile Met Tyr Met Ala Gly Asn Thr
                          -70
                                                                      100
atc tcc atc ttc cct act atg atg gtg tgt atg atg gcc tgg cga ccc
Ile Ser Ile Phe Pro Thr Met Met Val Cys Met Met Ala Trp Arg Pro
                   -55
                                        -50
                                                                      148
att cag gca ctt atg gcc att tca gcc act ttc aag atg tta gaa agt
Ile Gln Ala Leu Met Ala Ile Ser Ala Thr Phe Lys Met Leu Glu Ser
                                                        -30
               -40
                                    -35
                                                                      196
tca agc cag aag ttt ctt cag ggt ttg gtc tat ctc att ggg aac ctg
Ser Ser Gln Lys Phe Leu Gln Gly Leu Val Tyr Leu Ile Gly Asn Leu
                                                    -15
           -25
                                -20
atg ggt ttg gca ttg gct gtt tac aag tgc cag tcc atg gga ctg tta
Met Gly Leu Ala Leu Ala Val Tyr Lys Cys Gln Ser Met Gly Leu Leu
                            - 5
cct aca cat gca tcg gat tgg tta gcc ttc att gag ccc cct gag aga
                                                                      292
Pro Thr His Ala Ser Asp Trp Leu Ala Phe Ile Glu Pro Pro Glu Arg
                   10
                                        15
                                                                      340
atg gag tca gtg gtg gag gac tgc ttt tgt gaa cat gag aaa gca gcg
Met Glu Ser Val Val Glu Asp Cys Phe Cys Glu His Glu Lys Ala Ala
               25
                                    30
                                                                      388
cct ggt ccc tat gta ttt ggg tct tat tta cat cct tct tta agc cca
Pro Gly Pro Tyr Val Phe Gly Ser Tyr Leu His Pro Ser Leu Ser Pro
                                45
gtg gct cct cag cat act ctt aaa cta atc act tat gtt aaa aaa aac
Val Ala Pro Gln His Thr Leu Lys Leu Ile Thr Tyr Val Lys Lys Asn
                           60
                                                                      483
caa aaa act ctt ttc tcc atg gtg ggg tgacaggtcc taaaaggaca
Gln Lys Thr Leu Phe Ser Met Val Gly
                        75
                                                                      543
atgtgcatat tacgacaaac acaaaaaaac tataccataa cccagggctg aaaataatgt
                                                                      603
aaaaaacttt attttgttt ccagtacaga gcaaaacaac aacaaaaaaa cataactatg
                                                                      663
taaacaaaaa aataactgct gctaaatcaa aaactgttgc agcatctcct ttcaataaat
                                                                      696
taaatggttg araacaatgc aaaaaaaaaa aaa
```

```
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 263..481
<221> sig_peptide
<222> 263..322
<223> Von Heijne matrix
      score 11.1999998092651
      seq ILVVLMGLPLAQA/LD
<221> polyA site
<222> 858..868
<400> 263
aagacacgcc tacgattaga ctcaggcagg cacctaccgg cgagcggccg crvgtgactc
ccaggegegg eggtacetea eggtggtgaa ggtcaeaggg ttgcageaet eccagtagae
                                                                      120
caggagetee gggaggeagg geeggeeeca egteetetge geaceaceet gagttggate
                                                                      180
ctctgtgcgc cacccctgag ttggatccag ggctagctgc tgttgacctc cccactccca
                                                                      240
egetgeeete etgeetgeag ee atg acg eee etg ete ace etg ate etg gtg
                                                                      292
                         Met Thr Pro Leu Leu Thr Leu Ile Leu Val
                                             -15
                         -20
                                                                      340
gtc ctc atg ggc tta cct ctg gcc cag gcc ttg gac tgc cac gtg tgt
Val Leu Met Gly Leu Pro Leu Ala Gln Ala Leu Asp Cys His Val Cys
-10
gee tae aac gga gae aac tge tte aac eee atg ege tge eeg get atg
                                                                      388
Ala Tyr Asn Gly Asp Asn Cys Phe Asn Pro Met Arg Cys Pro Ala Met
                                15
            10
gtt gcc tac tgc atg acc acg cgc acc tac tac acc ccc acc agg atg
                                                                      436
Val Ala Tyr Cys Met Thr Thr Arg Thr Tyr Tyr Thr Pro Thr Arg Met
        25
                            30
                                                                      481
aag gtc agt aag tcc tgc gtg ccc cgc tgc ttc gar nac tgt gta
Lys Val Ser Lys Ser Cys Val Pro Arg Cys Phe Glu Xaa Cys Val
                        45
tgatggctac tccaagcacg cgtccaccac ctcctgctgc cagtacgacc tctgcaacgg
                                                                      541
                                                                      601
caceggeett gecacecegg ccacectgge cetggeeeec atecteetgg ccacectetg
gggteteete taaageeeee gaggeagace caeteaagaa caaagetete gagacacaet
                                                                      661
gctayaccct ckcacccake teaccetgee teacceteea caetecetge gaceteetea
                                                                      721
gccatgccca gggtcaggac tgtgggcaag aagacacccg acctccccca accaccacac
                                                                      781
gacctcactt cgaggccttg acctttcgat gctgtgtggg atcccaaaag tgtccggctt
                                                                      841
                                                                      868
tgatgggctg atcagcaaaa aaaaaaa
```

<211> 775

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 42..299

<221> sig_peptide

<222> 42..101

<223> Von Heijne matrix
 score 5.40000009536743
 seq WFVHSSALGLVLA/PP

<221> polyA_site

<222> 762..775

<400> 264 aacgatacaa atggtaggcc ttcatgtgag ccagtda	cta c atg aat ctt cat ttc 56  Met Asn Leu His Phe -20
cca cag tgg ttt gtt cat tca tca gcg tta Pro Gln Trp Phe Val His Ser Ser Ala Leu	ggc ttg gtc ctg gct cca 104
cct ttc tcc tct ccg ggc act gac ccc acc Pro Phe Ser Ser Pro Gly Thr Asp Pro Thr 5 10	ttt ccg tgt att tac tgt 152 Phe Pro Cys Ile Tyr Cys 15
agg cta tta aat atg atc atg acc cgc ctt Arg Leu Leu Asn Met Ile Met Thr Arg Leu 20 25	gca ttt tca ttc atc acc 200 Ala Phe Ser Phe Ile Thr 30
tgt tta tgc cca aat tta aag gaa gtt tgt Cys Leu Cys Pro Asn Leu Lys Glu Val Cys 35 40	ctc att ttg cca gaa aaa 248 Leu Ile Leu Pro Glu Lys 45
aat tgt aat agt cga cac gct gga ttt gta Asn Cys Asn Ser Arg His Ala Gly Phe Val	ggg cca sca aaa ttg cgg 296 Gly Pro Xaa Lys Leu Arg 60 65
cag tgaaactwkk ttcwcttcta aagcccttca ttt	.cccacaa ggttaagctc 349
togaaaccc atttgatcct tggttcctat ttcgatc tctccatgtt gtatgcaaat taaaakttgc cttgttt cagggaraaa gaggccttat ctgttcctcc atcccc ttcctcagga cttcctttgg ttggggattt tactttc tcaggggtag acaagcttgt cctagtgctc tgcttca aatagaaaag gtagatgcct tgacttttgt ccctgtt cagaattgtc aaaagctccg gttcaaactc tgtagagaaaaa	gtt actettecaa cacagggtat 469 eetg ttttgacaga etgetaagaa 529 eeca aaagtetgat etgatttett 589 eggt ettateagaa gaaacecagg 649 egtg gggactaaag tgttttttge 709

<210> 265
<211> 1075
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 198..431

<221> sig_peptide
<222> 198..260
<223> Von Heijne matrix
score 6.90000009536743
seq LLACGSLLPGLWQ/HL

<221> polyA_site <222> 1064..1074

<400> 265

atatattct gaggcagtac ccatctcact tgtaaactta aaagacaccg cagagatttg
agggactcag aagtcaaata gagtaggtta aaaacctctt atttttcaaa ttaattgttt
taagaaacaa gcatacctgt gtaagtgaaa tatcttaatt tgtgttgaat caagttagga
gacagagatt ctcatga atg tgt cct gtg ttc tca aag cag ctg cta gcc

Met Cys Pro Val Phe Ser Lys Gln Leu Leu Ala

-20

-15

tgt ggg tct ctc cta cct ggg tta tgg cag cac ctc aca gcc aat cac
Cys Gly Ser Leu Leu Pro Gly Leu Trp Gln His Leu Thr Ala Asn His
-10

-5

1

60

230

230

230

tgg cct cca ttc tcc sct ttc ctc tgt aca gtt tgc tct ggt tcc tca Trp Pro Pro Phe Ser Xaa Phe Leu Cys Thr Val Cys Ser Gly Ser Ser 10 15 20	326
gag cag att tcc gag tat act gct tca gcc acg ccc cca ctg tgc cgt Glu Gln Ile Ser Glu Tyr Thr Ala Ser Ala Thr Pro Pro Leu Cys Arg 25 30 35	374
tcc ctg aac caa gag cca ttc gty tca aga gcc att cgt cca aag tac Ser Leu Asn Gln Glu Pro Phe Val Ser Arg Ala Ile Arg Pro Lys Tyr 40 45 50	422
tct atc acc tagecattgt akccatacca ageegggett ectaetteee Ser Ile Thr 55	471
totgotocco tiggittoct cotginaari aaatotoaci gaccotigat goasotocaa	531
gcatatataa tatatata ataaaaccat abtctaaaaa attcaaacca ggawaaataa	591
asccaraaat ttgtatggga aaaatctgca caaatttatt tggccagcat ggttatcatg	651
gctctattga atttatcctt gaccgtcttt aaagccaaag caaacgggat aaagtgatca	711
actacttacc tctcaatacc aaaaargaag caggaggcaa aatctctcaw taatttcata	771
aaaacaatto ttakotgggo goggtggoto woacotgtar toocaacact ttgggaggoo saggtgggog gatoatgagg togggagato aamacoatoo tggotaacat ggtgaaacoo	831
catctctact aaaattacaa aaaattrgct gggcgaggtg gcgggcacct gtggtcccag	891 951
ctactcggga ggctgaggca agagaatggt gtgaacccca gggggcggag cctgcagtga	1011
gctgagatcg caccactgca ctccagcctg ggcgacagtg agactccgtc tcaaaaaaaa	1071
aaah	1075
<210> 266 <211> 981 <212> DNA <213> Homo sapiens	
<220> <221> CDS	
<222> CDS <222> 279473	
<221> sig_peptide	
<222> 279362	
<223> Von Heijne matrix	
score 4.40000009536743 seq SCFLVALIIWCYL/RE	
<221> polyA_signal <222> 944949	
<221> polyA_site <222> 970981	
<400> 266	
agaategtgt ettgtgtgee eeggeggeeg ggtgagetee teaaggtete ggagggeega gggeagaeae eggegggegg geggasgett aetgetetet etetteeagg geegteeggg	60 120
egetgagget cataggetgg getteeegaa geetteatee gttgeeeggt teeegggate	120 180
gggcccaccc tgccgccgag gaagaggacg accetgaccg ccccattgag ttttcctcca	240
gcaaagccaa ccctcaccgc tggtcggtgg gccatacc atg gga aag gga cat cag	296
Met Gly Lys Gly His Gln -25	
CGG CCC tGG tGG aag GtG CtG CCC CtC agC tGC ttC CtC GtG GCG CtG  Arg Pro Trp Trp Lys Val Leu Pro Leu Ser Cys Phe Leu Val Ala Leu  -20 -15 -10	344
atc atc tgg tgc tac ctg agg gag gag agc gag gcg gac cag tgg ttg	392
Ile Ile Trp Cys Tyr Leu Arg Glu Glu Ser Glu Ala Asp Gln Trp Leu	
-5 1 5 10	
aga cag gtg tgg gga gag gtg cca gag ccc agt gat cgt tct gag gag	440

Arg Gln Val Trp Gly Glu Val Pro Glu Pro Ser Asp Arg Ser Glu Glu	
15 20 25	
cct gag act cca gct gcc tac aga gcg aga act tgacggggtg cccgctgggg.  Pro Glu Thr Pro Ala Ala Tyr Arg Ala Arg Thr  30 35	493
ctggcaggaa gggagccgac asccgccctt cggatttgat ktcacgtttg cccgtgactg	553 613
tootggotat goktgogtoo toagoaotra argaottggo tggtggatgg ggcaottggo tatgotgatt ogogtgaagg oggavoaaaa totoagoaaa toggaaactg otootosoot	673
ggctcttgat ktccaaggat tccatcggca aaacttctca ratccttggg gaaggtttca	733
gttgcactgt atgctgttgg atttgccaag tctttgtata acataatcat gtttccaaag	793
cacttetggt gacacttgte atceagtgtt agtttgcagg taatttgctt tetgagatag	853 913
aatatotggo agaagtgtga aactgtattg catgotgogg cotgtgoaag gaacaottoo acatgtgagt tttacacaac aacaaatgaa aataaatttt aattttataa tatgggaaaa	973
aaaaaaaa	981
<210> 267	
<211> 1031	
<212> DNA <213> Homo sapiens	
22137 HOMO Sapiens	
<220>	
<221> CDS <222> 12644	
<222> 12644	
<221> sig_peptide	
<222> 1292 <223> Von Heijne matrix	
score 4	
seq LTFFSGVYGTCIG/AT	
<221> polyA signal	
<222> 10021007	
cally maly mains	
<221> polyA_site <222> 10201031	
<pre>&lt;400&gt; 267 acaccaagga g atg ctc ctt ctt agt att aca act gct tat aca ggt ctg</pre>	50
Met Leu Leu Ser Ile Thr Thr Ala Tyr Thr Gly Leu	
-25 -20 -15	98
gaa tta act ttc ttc tct ggt gta tat gga acc tgt att ggt gct aca Glu Leu Thr Phe Phe Ser Gly Val Tyr Gly Thr Cys Ile Gly Ala Thr	90
-10 -5 1	
aat aaa ttt gga gca gaa gag ara agc ctt att gga ctt tct ggc att	146
Asn Lys Phe Gly Ala Glu Glu Xaa Ser Leu Ile Gly Leu Ser Gly Ile 5 10 15	
5 10 15 ttc atc ggc att gga gaa att tta ggt gga agc ctc ttc ggc ctg ctg	194
Phe Ile Gly Ile Gly Glu Ile Leu Gly Gly Ser Leu Phe Gly Leu Leu	
20 25 30	242
agc aag aac aat cgt ttt ggt aga aat cca gtt gtg ctg ttg ggc atc Ser Lys Asn Asn Arg Phe Gly Arg Asn Pro Val Val Leu Leu Gly Ile	242
35 40 45 50	
ctg gtg cac ttc ata gct ttt tat cta ata ttt ctc aac atg cct gga	290
Leu Val His Phe Ile Ala Phe Tyr Leu Ile Phe Leu Asn Met Pro Gly	
55 60 65 gat gcc ccg att gct cct gtt aaa gga act gac agc agt gct tac atc	
	<b>33</b> 8
	338
Asp Ala Pro Ile Ala Pro Val Lys Gly Thr Asp Ser Ser Ala Tyr Ile 70 75 80	338
Asp Ala Pro Ile Ala Pro Val Lys Gly Thr Asp Ser Ser Ala Tyr Ile	338

85 90 95	
gga aac agc tgc ttt aat acc cas ctg ctt akt atc tkg ggc ttt ctg	434
Gly Asn Ser Cys Phe Asn Thr Xaa Leu Leu Xaa Ile Xaa Gly Phe Leu	
100 105 110 tat tot gaa rac ago goo coa koa ttt goo ato tto aat ttt gtt cag	482
Tyr Ser Glu Xaa Ser Ala Pro Xaa Phe Ala Ile Phe Asn Phe Val Gln	
115 120 125 130	
tet att tge gea gee gtg gea ttt tte tae age aac tae ett ete ett	530
Ser Ile Cys Ala Ala Val Ala Phe Phe Tyr Ser Asn Tyr Leu Leu Leu 135 140 145	
cac tgg caa ctc ctg gtc atg gtk atw ttt ggg ttt ttk gga aca att	578
His Trp Gln Leu Leu Val Met Val Ile Phe Gly Phe Xaa Gly Thr Ile	
150 155 160	
tot tto tto act gtg gaa tgg gaa sot goo goo ttt gta soc cgc ggc	626
Ser Phe Phe Thr Val Glu Trp Glu Xaa Ala Ala Phe Val Xaa Arg Gly 165 170 175	
tot gao tac oga agt ato tgatotggtg toogtgaggg gacacgtatg	674
Ser Asp Tyr Arg Ser Ile	
180	734
acctcagaaa cacagctgga cacagagctt ggtggaagaa gtcgcctttg atcttcacta tatattgggt gatgttcagt atggaaaatc aagggattaa gactgttaaa tcagccagag	794
tkggtgttca agtttacaga tatgagttat ttaaagcaag tagaataagg gaaagctgtt	854
ctgtcaactg taattgttca aagatgttgt ttttcatttc atctatctca attcttataa	914
tcatgttata gaatgtaaat gttttcttct ctctcctgct cttgttggaa gatcctgcct	974
tgatttagaa tactaggcca tatgtcatat aaatattttt tctggaaaaa aaaaaaa	1031
<pre>&lt;210&gt; 268 &lt;211&gt; 1283 &lt;212&gt; DNA &lt;213&gt; Homo sapiens  &lt;220&gt; &lt;221&gt; CDS &lt;222&gt; 91.459  &lt;221&gt; sig_peptide &lt;222&gt; 91.330 &lt;223&gt; Von Heijne matrix</pre>	
tattccttgg agttccacga ctgaattaag actgttgtgg grdccataat tttcaaatac	60
ttgccctata ttcgtgttga gggttcacac atg agc aca tgg tat ttg gca ctt	114
Met Ser Thr Trp Tyr Leu Ala Leu -80 -75	
aat aag too tat aag aat aaa gac ago gtt agg att tat oto ago ttg	162
Asn Lys Ser Tyr Lys Asn Lys Asp Ser Val Arg Ile Tyr Leu Ser Leu	
-70 -65 -60	
tgc aca gtg agc att aaa ttt aca tac ttt cat gat ata cag act aat Cys Thr Val Ser Ile Lys Phe Thr Tyr Phe His Asp Ile Gln Thr Asn	210
-55 -50 -45	
tgt ctt aca aca tgg aaa cat tcg aga tgc aga ttt tat tgg gca ttt	258
Cys Leu Thr Thr Trp Lys His Ser Arg Cys Arg Phe Tyr Trp Ala Phe	
-40 -35 -30 -25 ggt ggt tee att tta cag cae tea gtg gat eee ett gtt ttg tte eta	306
Gly Gly Ser Ile Leu Gln His Ser Val Asp Pro Leu Val Leu Phe Leu	500

•	
-20 -15 -10	
age etg gee etg tta gtg aca eee act tee ace eet tet get aar ata	354.
Ser Leu Ala Leu Leu Val Thr Pro Thr Ser Thr Pro Ser Ala Lys Ile	
-5 1 5	
car ago ott caa att gao oto oot gga ggo tgg agg otg goo act gao	402
Gln Ser Leu Gln Ile Asp Leu Pro Gly Gly Trp Arg Leu Ala Thr Asp	
10 15 20	
agg atc ttt acc ctc tcc ccc gta ccc atg gac rgc ccc ctc atc ctt	450
Arg Ile Phe Thr Leu Ser Pro Val Pro Met Asp Xaa Pro Leu Ile Leu	
25 30	
cat cag ttg taaaggtaga tatttgttcc ttggagtcca acatcatgct	499
His Gln Leu	
gttcagaata taatgagatc aatagttgaa aaactagata tacatgccac ccwgacaaag	559
ctattaagtt attaagtgtc agccctggat cttggcttat tgtgaaatgt taattatttt	619
atcactcyat taagaagctg tgggctccat ctcagcattg aaaagggact aatttgctct	679
gttttggaat tgaattaget tteaggeeas eagggeactg tttggtaaat tgetttttee	739
agtactagca tgttttctcc ctccatagcc tctgttagct tctgagcttg taacctccag	799
ggaaavatga gaatattcac ccttttaata tgtgtagaga ccatgcaaga ccattgtctt	859
ctaataatta gaaatactta gccagattct ctatagtaaa cccggagatt gggagggctg	919
ctttctactt ggtgcatcct tctgcgcttc taatgatttt taaaaatctg ttaataattg	979
atgttttctg gctgggcaca gtggctcacg cctgtaatcc cagcactttg ggaggccaag	1039
gagggcagat catgaggtca ggagattgar accatectgg ctaacacggt gaaaccccgt	1099
ctctactaaa aatacaaaar aattakccgg gcatggtagt gggcgcctgt gtacccagct	1159
actggggagg ctgaggcarg araatcgctt gaacctggga ggcggaggtt gcastragct	1219 1279
gagatggtgc caccgcactc tagcctgggt gacagagcga gacttcattt caaaaaaaaa	
aamc	1283
<210> 269	
<211> 1777	
\4±17 ±1//	

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 70..327

<221> sig_peptide

<222> 70..147 <223> Von Heijne matrix score 9.60000038146973 seq WLIALASWSWALC/RI

<221> polyA_signal <222> 1741..1746

<221> polyA_site <222> 1763..1774

<pre>&lt;400&gt; 269 agcccggttt cgtgc agagctaca atg ga Met Gl</pre>	a aag too tgg a	atg ctg tgg aad		a tgg 111
-2	25	-20	-15	
cta ata gcc ttg				
Leu Ile Ala Leu		r Trp Ala Leu (	Cys Arg Ile Ser	Leu
-10	-5		1	
tta cct tta ata	gtg act ttt cat	t ctg tat gga 🤉	ggc att atc tta	ctt 207
Leu Pro Leu Ile	Val Thr Phe His	s Leu Tyr Gly	Gly Ile Ile Leu	Leu
5	10	15		20
ttg tta ata ttc	ata tca atw kc	a ggt att ctg	tat aaa ttc cas	gat 255

	•
Leu Leu Ile Phe Ile Ser Ile Xaa Gly Ile Leu Tyr Lys Phe Xaa Asp	
Leu Leu He Phe He Ser He Kaa Giy He Leu Tyr Lys The Maa 125	
gta ttg ctt tat ttt ccw kaa cag yya tcc tct tca cgt ctt tat gat	303
Val Leu Leu Tyr Phe Pro Xaa Gln Xaa Ser Ser Ser Arg Leu Tyr Asp	
40 45 50	
tcc cat gcc cac tgg cmt tcg rca taaaaaaatt ttcatcagaa ccaaagatgg	357
Ser His Ala His Trp Xaa Ser Xaa	
55 60	
aatacgtctg aatcttattt tgatacgata cactggagac aattcaccct attccccaac	417
tataatttat tttcatggga atgcaggcaa cataggtcac aggttggcca aatgcattac	477
ttatgttggt taacctcaaa gttaaccttt tgctggttga ttatcgagga tatggaaaaa	537
gtgaaggaga agcaagtgaa gaaggactct acttagattc tgaagctgtg ttagactacg	597
tgatgactag acctgacctt gataaaacaa aaatttttct ttttggccgt tccttgggtg	657
garcagtggc tattcatttg gcttctgaaa attcacatag gatttcagcc attatggtgg	717
agaacacatt tttaagcata ccacatatgg ccagcacttt attttcattc tttccgatgc	777
gttaccttcc tttatggtgc tacaaaaata aatttttgtc ctacagaaaa atctctcagt	837 897
gtagaatgcc ttcacttttc atctctggac tctcagatca attaattcca ccagtaatga	957
tgaaacaact ttatgaactc tccccatctc ggactaagan attagccatt tttccagatg	1017
ggactcacaa tgacacatgg cagtgccaag gctatttcac tgcacttgaa cagttcatca	1017
aagaagtogt aaagagcoat totootgaag aaatggcaaa aacttoatot aatgtaacaa	1137
ttatataatg tttccctttt tgattattgc attgtatttt aatttgtgca gaatgataaa	1197
gaatgttcct tttagaagtg tgttatgtct gtacctgtct gaagagtgac attaaacttt	1257
gaaaggactt cactgctcct ttacgatatt ccaaatagtt ttttacattg gaaaaactaa	1317
ttottgggat totttcatac attttcatca aaactttcag tgtgattatg tattcatatc	1377
ttcagtttaa tatgtcagta taatagatat tgttcaaaag tttcttgttg ctaaagtggt	1437
gtaatctgtt acacagatga atagctagat gtggaaagag atatgtaaac aagaaacctt	1497
tgggtattgt ttcttaagta aatattggga caatcatggt aagcaaactt agttctgtaa	1557
ctgcattttt caccttaaaa gttaaatgaa atgcatgatg gtattttatt ccttgaatta	1617
tgcaatgcaa cattttacat gtaaatagca ctggtcatat actgatgtat atggttatct gggttatatc tatttttatg taaactctat ttttgttttt ggcaagaagt gaaattgaga	1677
cttatgtgca ggttgccatt gaattttgct ctggtgaatg ctgagatcca gctttttctt	1737
acaaataaat gggaccctgt tttccaaaaa aaaaaaamcm	1777
acaaataaat gggaccctgt tttccaaaaa aaaaaaamcm	
<210> 270	
<211> 970	
<211> 570 <212> DNA	
<213> Homo sapiens	
•	
<220>	
<221> CDS	
2225 12 407	

			-15					-10					-5			
aca	gga	ccc	tgg	999	gct	gtt	gcc	acc	tcc	gcc	999	ggc	gag	gag	tcg	146
Thr	Gly	Pro	Trp	Gly	Ala	Val	Ala	Thr	Ser	Ala		Gly	Glu	Glu	Ser	
		1				5					10			~~t	663	194
ctt	aag	tgc	gag	gac	ctc	aaa	gtg	gga	caa	Tat	Tla	Cyc	Lac	gat	Dro	134
ьеи 15	гув	Cys	GIU	Asp	20	гур	vai	GIY	GIII	25	116	Cys	טעט	Asp	30	
	ata	aat	gac	act		caa	gaa	cca	att		tat	aca	aac	tac	aca	242
Lvs	Ile	Asn	Asp	Ala	Thr	Gln	Glu	Pro	Val	Asn	Cys	Thr	Asn	Tyr	Thr	
-			-	35					40					45		
gct	cat	gtt	tcc	tgt	ttt	cca	gca	ccc	aac	ata	act	tgt	aag	gat	tcc	290
Ala	His	Val		Cys	Phe	Pro	Ala		Asn	Ile	Thr	Cys		Asp	Ser	
			50					55	224	~~~	~++	cat	60	++~	220	338
agt	ggc	aat	gaa	aca Thr	Tic	Dhe	Thr	999 G1v	Aen	Glu	Val	Glv	Phe	ttc Phe	Lvs	330
261	Gry	65	GIU	1111	1114	riic	70	OL J		014		75			-1 -	
ccc	ata		tgc	cga	aat	gta	aat	ggc	tat	tcc	tac	aat	gag	cag	tcg	386
Pro	Ile	Ser	Cys	Arg	Asn	Val	Asn	Gly	Tyr	Ser	Tyr	Asn	Glu	Gln	Ser	
	80					85					90					
cat	gtc	tct	ttt	tct	tgg	atg	gtt	<b>a</b> aa	agc	aga	tcg	att	tta	cct	tgg	434
	Val	Ser	Phe	Ser	Trp	Met	vai	GIĀ	Ser	Arg	ser	TIE	ьеи	Pro	11p	
95 25 2	666	tac	+++	aaa		att	222	htt	tva		tat	agg	att	tkg		482
Tle	Pro	Cvs	Phe	61 A	Phe	Val	Lvs	Xaa	Xaa	His	Cvs	Arq	Val	Xaa	Trp	
		<i>u                                    </i>		115			2		120		-			125	-	
aat	tgg	gag	cct	aat	tgai	tttc	aty (	cttai	ttc	aa t	gcaga	attg	t tg	gacci	ttca	537
Asn	Trp	Glu		Asn												
			130										act /	aaat:	attact	597
aat	ggaa	gta (	gtta	catt.	at ag	gatt	tata	t gga	aacc	agac	tttt	traa	acc y	aaac	attact agattt	657
															ttttc	717
att	cata	tac	catt	ttat	ga gi	ttct	gtat	a ati	tttt	tgtg	gtti	tttg	ttt	tgtt	gagtta	777
aag	tata	tta '	ttgt	gaga	tt t	attt	aata	g ga	cttc	cttt	gaaa	agct	gta '	taat	agtgtt	837
tct	cggg	ctt	ctgt	ctct	at g	agag	atag	c tta	atta	ctct	gata	actc	ttt i	aatc	ttttac	897
aaa	ggca	agt '	tgcc	actt	gt c	attt	ttgt	t tc	tgaa	aaat	aaaa	agta	taa	cttai	ttcaca	957
aaa	aaaa	aaa 1	mms													970

```
<210> 271
```

<220>

<221> polyA_signal

<211> 645

<212> DNA

<213> Homo sapiens

<221> CDS

<222> 90..383

<221> sig_peptide <222> 90..200

<223> Von Heijne matrix score 4.90000009536743 seq MLIMLGIFFNVHS/AV

<222> 609..614

<221> polyA_site

<222> 632..643

WO 9	9/3123	6						-1	93-			•		PCT/IE	98/02122
cacctcg							Met	Ala	-35	Leu	Leu	Cys	Cys.	-30	113
ccg aag Pro Lys	ctg Leu	gcc Ala	gcc Ala -25	tgc Cys	ggc	atc Ile	gtc Val	ctc Leu -20	agc Ser	gcc Ala	tgg Trp	gga Gly	gtg Val -15	atc Ile	161
atg ttg Met Leu	ata Ile	atg Met -10	ctc	gga Gly	ata Ile	ttt Phe	ttc Phe -5	aat Asn	gtc Val	cat His	tcc Ser	gct Ala 1	gtg Val	ttg Leu	209
att gag	gac Asp	att	ccc Pro	ttc Phe	acg Thr 10	gag Glu	aaa Lys	gat Asp	ttt Phe	gag Glu 15	aac Asn	ggc	ccc Pro	car Gln	257
aac ata Asn Ile	tac Tyr	aac Asn	ctt Leu	tac Tyr 25	rag	caa Gln	ktc Xaa	agc Ser	tac Tyr 30	aac Asn	tgt Cys	ttc Phe	atc Ile	gct Ala 35	305
gca ggc Ala Gl	ctt Leu	tac Tyr	Leu	ctc	ctc Leu	gga Gly	ggc Gly	ttc Phe 45	tct	ttc Phe	tgc Cys	caa Gln	ktt Xaa 50	cgg Arg	353
ctc aat Leu Asi	aag n Lys	Arg	40 aag Lys	gaa Glu	tac Tyr	atg Met	gtg Val 60	cgc	tag	ggcc	ccg	gcgc	-	cc	403
ccgctc		55 ccct	cctc	ta t	ttaa	arac	t. cc	ctgc	accg	tkt	cacc	cag	gtcg	cgtccc	463
accette	rcca	acac	coto	ta t	aaaa	ctaa	q tt	tccc	gggc	rar	arac	tga	acco		523
ccatct	ataa	cato	caac	cc c	cata	qara	r qq	ctga	ggcu	999	9990	Lg L	cccg		583
caccct	tcgc	tgtg	tccc	gt a	tctc	aata	a ag	agaa	tctg	ctc	tctt	caa	aaaa	aaaaaa	643 645
<210> <211> <212> <213> <213> <220> <221> <222> <222> <222> <222>	773 DNA HOMO CDS 332. sig_]	.541 pept: .376 Heij: e 3.9	ide ne ma	99904	6325	57									
<221> <222> <221> <222>	739.	.744 A_si													
<400>	272				~-	++ = <del>+</del>	ta t	tatt	aaad.	t of:	aaac	aaaa	tta	caatttc	60
22222		2 F C	~~~~		act C				~~~~	1			9		

aaaacaattc atgcctttca tagtttatta ttattaaagt ctaaacaaaa ttgcaatttc ttaggtaacc ttatatttac aataaatgaa gattaccctc aaatgctaga agctgtctag 120 gtccgtccgg tgtgtcagat tttcctcaga ttagatgtgc caataaccaa gtttattcag 180 taaacaactt gtacttgttt catctggttt tattactctc acccataaac agtaatgact 240 ctctgaccct ctggaaatat gtaatgcttc caatcttgct ttgtgtatct catttaattt 300 gttataaggt agtactgatt ttagcatatt a atg cga ttt ctt cct tgt tgt 352 Met Arg Phe Leu Pro Cys Cys -10 -15 ttg ctt tgg tct gtg ttc aat cca gag agc tta aat tgt cat tat ttt 400 Leu Leu Trp Ser Val Phe Asn Pro Glu Ser Leu Asn Cys His Tyr Phe -5

	ndd	gaa	a m.c	tat	a++	+++	on t		tt 2				gaa	att	tca	448
Xaa 2	•		aiiic		aLL		gyc	agt	CCA	caa	tat	tat	9~~			
	(aa	Glu	Xaa	Сув	Ile	Phe	Xaa	Ser	Leu	Gln	Tyr	Tyr	Glu	Ile	Ser	
	LO			•		15					20	•		•		
tt d	aa	gag	aaa	cta	cta	qqc	ttc	ctg	tgg	ctt	tgt	ttt	ctt	agt	tac	496
eu (	3ln	Glu	Lvs	Leu	Leu	Gly	Phe	Leu	Trp	Leu	Cys	Phe	Leu	Ser	Tyr	
5			-3-		30	•			-	35	-				40	
.++ 1	-+-	cat	acc	ata		ttt	tta	att	gat	ttt	tat	tct	ttt	act		541
he i	Dhe	Ara	Δla	Val	TVT	Phe	Leu	Ile	Asp	Phe	Ser	Ser	Phe	Thr		
riie .	110	9		45	-1-				50					55		
	2272	aa c	tatt		+ ++	caaa	atoto	ato	cata	ttt	acat	tcta	agt '	tcaga	agccaa	601
·gaa		oct c	,-3	raat	t to	cact	gtaa	tta	aaaac	tat	ttac	atatt	tag	ttata	aaatag	661
3000			, , , , , ,	ratto	·+ cc	atta	caco	ato	acct	gca	tcac	age	ca	taato	gaatgt	721
+~+		.ac	tage	rass	t as	aaat	gaca	aat	gcac	tga	aaaa	aaaa	aaa	aa		773
<210	> 2'	73														
<211	> 56	56														
<212																
			sapie	ens												
			•													
<220	>															
<221		os														
<222	> 4	322	22													
<221	> S	ig_pe	eptic	ìe												
		31														
<223	> V	on He	eijne	e mat	crix											
	S	core	4													
	s	eq El	NFLSI	LLSK	SCSA,	/DP										
		-														
<221			_sigr													
	q <		_sigr													
	q <	olyA	_sigr													
<222 <221	> p > 5 > p	olyA 30	_sigr 535 _site	nal												
<222 <221	> p > 5 > p	olyA 30	_sigr 535 _site	nal												
<222 <221 <222	> p > 5 > p > 5	olyA 30	_sigr 535 _site	nal												
<222 <221 <222 <400	> p > 5 > p > 5 > 2	olyA 30 olyA 55	_sign 535 _site 566	nal									<b>.</b>	299	c.t.t	5.
<222 <221 <222 <400 aacg	> p > 5 > p > 5 > 2	olyA 30 olyA 55 73 gga	_sign 535 _site 566 ggtg	nal e							1	Met : -45	His	agc Ser	Leu	5
<222 <221 <222 <400 aacg	> p > 5 > p > 5 > 2 agt	olyA 30 olyA 55 73 gga	_sign 535 _site 566 ggtg	nal e tggc	aaa	gtt	ctt	ttc	tat	tac	agt	Met : -45 ttt	His ago	ser ttt	ьец agg	
<222 <221 <222 <400 aacg	> p > 5 > p > 5 > 2 agt	olyA 30 olyA 55 73 gga	_sign 535 _site 566 ggtg	nal e tggc	aaa	gtt	ctt	ttc	tat	tac	agt	Met : -45 ttt	His ago	ser ttt	ьец agg	
<222 <221 <222 <400 aacg ttc Phe	> p > 5 > p > 5 > 2 agt atte -40	olyA 30 olyA 55 73 gga ggg Ala	_sign 535 _site 566 ggtg	tggc ttg	aaa Lys	gtt Val -35	ctt Leu	ttc Phe	tat Tyr	tac Tyr	agt Ser	Met -45 ttt Phe	His ago Ser	ser ttt	agg Arg	10
<222 <221 <222 <400 aacg ttc Phe	> p > 5 > 5 > 2 agt atle aat	olyA 30 olyA 55 73 gga ggga	_sign 535 _site 566 ggtge agc Ser	tggc ttg Leu	aaa Lys tgc	gtt Val -35 ctt	ctt Leu ctc	ttc Phe cac	tat Tyr aat	tac Tyr	agt Ser -30	-45 ttt Phe	ago Ser aat	ser ttt Phe	agg Arg ctt	5. 10: 15
<222 <221 <222 <400 aacg ttc Phe	> p > 5 > 5 > 2 agt atle aat	olyA 30 olyA 55 73 gga ggga	_sign 535 _site 566 ggtge agc Ser	tggc ttg Leu	aaa Lys tgc	gtt Val -35 ctt	ctt Leu ctc	ttc Phe cac	tat Tyr aat	tac Tyr	agt Ser -30	-45 ttt Phe	ago Ser aat	ser ttt Phe	agg Arg ctt	10
<222 <221 <222 <400 aacg ttc Phe ttt Phe -25	> p p 5 > 5 > 2 agt atte -40 aatt Asn	olyA 30 olyA 55 73 gga ggg Ala tgg	_sign 535 _site 566 ggtg agc Ser ttc Phe	tggc ttg Leu gac Asp	aaa Lys tgc Cys -20	gtt Val -35 ctt Leu	ctt Leu ctc Leu	ttc Phe cac His	tat Tyr aat Asn	tac Tyr ttg Leu	agt Ser -30 ggc Gly	Met -45 ttt Phe gag Glu	agc Ser aat Asr	ser ttt Phe ttc	agg Arg ctt Leu -10	10.
<222 <221 <222 <400 aacg ttc Phe ttt Phe -25 agc	>> 5 p 5 s agt atted Asn	olyA 30 olyA 55 73 gga ggg Ala tgg Trp	_sign 535 _site 566 ggtg agc Ser ttc Phe	tggc ttggc Leu gac Asp	aaa Lys tgc Cys -20	gtt Val -35 ctt Leu	ctt Leu ctc Leu	ttc Phe cac His	tat Tyr aat Asn	tac Tyr ttg Leu -15	agt Ser -30 ggc Gly	Met -45 ttt Phe gag Glu	ago Ser aat Asr	ttt Phe ttc Phe act	agg Arg ctt Leu -10 ttc	10:
<222 <221 <222 <400 aacg ttc Phe ttt Phe -25 agc	>> 5 p 5 s agt atted Asn	olyA 30 olyA 55 73 gga ggg Ala tgg Trp	_sign 535 _site 566 ggtg agc Ser ttc Phe	tggc ttggc Leu gac Asp	aaa Lys tgc Cys -20	gtt Val -35 ctt Leu	ctt Leu ctc Leu	ttc Phe cac His	tat Tyr aat Asn	tac Tyr ttg Leu -15	agt Ser -30 ggc Gly	Met -45 ttt Phe gag Glu	ago Ser aat Asr	ttt Phe ttc Phe act	agg Arg ctt Leu -10 ttc	10:
<222 <221 <222 <400 aacg ttc Phe ttt Phe -25 agc	>> 5 p 5 s agt atted Asn	olyA 30 olyA 55 73 gga ggg Ala tgg Trp	_sign 535 _site 566 ggtg agc Ser ttc Phe	tggc ttg Leu gac Asp aaa Lys	aaa Lys tgc Cys -20	gtt Val -35 ctt Leu	ctt Leu ctc Leu	ttc Phe cac His	tat Tyr aat Asn	tac Tyr ttg Leu -15	agt Ser -30 ggc Gly	Met -45 ttt Phe gag Glu	ago Ser aat Asr	ttt Phe ttc Phe act	agg Arg ctt Leu -10	10:
<2222 <221 <222 <400 aacg ttc Phe ttt Phe -25 agc Ser	> p > 5 > p > 2 agt atte -40 aatt Leu	olyA 30 olyA 55 73 gga gcg Ala tgg Trp ctc Leu	_sign 535 _site 566 ggtg agc Ser ttc Phe	tggc ttggc Leu gac Asp aaa Lys	aaa Lys tgc Cys -20 agt Ser	gtt Val -35 ctt Leu tgt	ctt Leu ctc Leu tct	ttc Phe cac His gcg Ala	tat Tyr aat Asn gac Asp	tac Tyr ttg Leu -15 ccg Pro	agt Ser -30 ggc Gly tct Ser	Met -45 ttt Phe gag Glu ggg Gly	ago Ser aat Asr tca Ser 5	ttt Phe ttc Phe act	agg Arg ctt Leu -10 ttc	10:
<222 <221 <222 <400 aacg ttc Phe ttt Phe -25 agc Ser atg	> p > 5 > p > 5 > 2 agt attle -40 aatt Leu agg	olyA 30 olyA 55 73 gga gcg Ala tgg Trp ctc Leu gac	_sign 535 _site 566 ggtg agc Ser ttc Phe agc	tggc ttg Leu gac Asp aaa Lys gag	aaa Lys tgc Cys -20 agt Ser	gtt Val -35 ctt Leu tgt Cys	ctt Leu ctc Leu tct Ser	ttc Phe cac His gcg Ala	tat Tyr aat Asn gac Asp	tac Tyr ttg Leu -15 ccg Pro	agt Ser -30 ggc Gly tct Ser	Met -45 ttt Phe gag Glu ggg Gly	ago Ser aat Asr tca Ser 5	ttt Phe ttc Phe act	agg Arg ctt Leu -10 ttc	10: 15
<222 <221 <222 <400 aacg ttc Phe ttt Phe -25 agc Ser atg	> p > 5 > p > 5 > 2 agt attle -40 aatt Leu agg	olyA 30 olyA 55 73 gga gcg Ala tgg Trp ctc Leu gac Asp	_sign 535 _site 566 ggtg agc Ser ttc Phe	tggc ttg Leu gac Asp aaa Lys gag	aaa Lys tgc Cys -20 agt Ser	gtt Val -35 ctt Leu tgt Cys	ctt Leu ctc Leu tct Ser	ttc Phe cac His gcg Ala	tat Tyr aat Asn gac Asp	tac Tyr ttg Leu -15 ccg Pro	agt Ser -30 ggc Gly tct Ser	Met -45 ttt Phe gag Glu ggg Gly	ago Ser aat Asr tca Ser 5	ttt Phe ttc Phe act	agg Arg ctt Leu -10 ttc	10 15
<222 <221 <222 <400 aacg ttc Phe ttt Phe -25 ager atg Met	> p > 5 > p > 5 > 2 agt attle -40 aatt Asn ctt Leu agg	olyA 30 olyA 55 73 gga gcg Ala tgg Trp ctc Leu gac Asp	_sign 535 _sitc 566 ggtg agc Ser ttc Phe agc Ser att	tggc ttg Leu gac Asp aaa Lys gag Glu	aaa Lys tgc Cys -20 agt Ser aca	gtt Val -35 ctt Leu tgt Cys aac Asn	ctt Leu ctc Leu tct Ser aaa Lys	ttc Phe cac His gcg Ala	tat Tyr aat Asn gac Asp 1	tac Tyr ttg Leu -15 ccg Pro	agt Ser -30 ggc Gly tct Ser	Met -45 ttt Phe gag Glu 999 Gly	ago Ser aat Asr Ser 5	ttt Phe ttc Phe act Thr	agg Arg ctt Leu -10 ttc Phe	10: 15
<222 <221 <222 <400 aacc Phe tte -25 ager atg Met	> 5 P P > 5 P P P P P P P P P P P P P P	olyA 30 olyA 55 73 gga gcg Ala tgg Trp ctc Leu gac Asp 10	_sign 535 _sitc 566 ggtg agcr ttc Phe agcr ser tle arac	tggc ttgu Leu gac Asp aaa Lys gagu	aaa Lys tgc Cys -20 agt Ser aca Thr	gtt Val -35 ctt Leu tgt Cys aac Asn	ctt Leu ctc Leu tct Ser aaa Lys gctg	ttc Phe cac His gcg Ala tga	tat Tyr aat Asn gac Asp 1 aata	tac Tyr ttg Leu -15 ccg Pro tgg	agt Ser -30 ggc Gly tct Ser gtta	Met -45 ttt Phe gag Glu ggg Gly aagt	Asr. Ser Ser Ser Ser Ser Ser	tttt Phe ttc Phe Thr ctga	agg Arg ctt Leu -10 ttc Phe gcagc	10: 15 19
<222 <221 <222 <400 aacc Phe tte -25 ager atg Met taca	> 5 5 7 5 7 5 7 7 7 7 7 7 7 7 7 7 7 7 7	olyA 30 olyA 55 73 gga ggg Ala tgg Trp ctc Leu gac Asp 10 aga	_sign 535 _sitc 566 ggtg agcr ttc Phe agcr atle arac	tggc ttgu Leu gasp aaa Lys galu cagt	aaa Lys tgc Cys -20 agt Ser aca Thr	gtt Val -35 ctt Leu tgt Cys aac Asn	ctt Leu ctc Leu tct Ser aaa Lys gctg	ttc Phe cac His gcg Ala tga g ag	tat Tyr aat Asn gac Asp 1 aata	tac Tyr ttg Leu -15 ccg Pro tgg	agt Ser -30 ggc Gly tct Ser gtta cac	Met -45 ttt Phe gag Glu ggg Gly aagt	Asr. Ser Ser Ser Asr. Ser Ser	ser tttt Phe ttc Phe Thr ctga aaga gcat	agg Arg ctt Leu -10 ttc Phe gcagc	10: 15 19 25
<222 <221 <222 <400 aacc Phe tte -25 ager atg tec tgcag	> 5 5 7 5 7 5 7 5 7 5 7 7 5 7 7 7 7 7 7	olyA 30 olyA 55 73 gga  gcg Ala tgg Trp ctc Leu gac Asp 10 aga aga	_sign 535 _sitc 566 ggtgccr Ser the ager alle arac tgaa	tggcttuucachaal	aaa Lys tgc Cys -20 agt Ser aca Thr	gtt Val -35 ctt Leu tgt Cys aac Asn tcct	ctt Leu ctc Leu tct Ser aaa Lys gctg agag cara	ttc Phe cac His gcg Ala tga g cc c tg	tat Tyr aat Asn gac Asp 1 aata	tac Tyr ttg Leu -15 ccg Pro tgg	agt Ser -30 ggc Gly tct Ser gtta cac aaa	Met -45 ttt Phe gag Glu ggg Gly aagt acag tcag	Asr ac to ara	ser ttt Phe ttc Phe ttc Thr ctga aaga gcat cttg	agg Arg ctt Leu -10 ttc Phe gcagc	10: 15 19 25 31 37
<222 <221 <222 <400 aa tte the tte-25c abet tacag tteas	>> 5 P P S S S S S S S S S S S S S S S S S	olyA 30 olyA 55 73 gga Ala tggp Trp cteu gac Asp 10 aga agaa	_sign 535 _sitc 566 gg agr ser ce Phe agr alle arac argaq tgaq	tggc ttu gasp aas sgalu cagaaaagagaga	aaa Lys tgc Cys -20 agt Ser athr ct gc ttc	gtt Val -35 ctt Leu tgt Cys aac Asn tcct aaag	ctt Leu ctc Leu tct Ser aaa Lys gctg agag cara	ttc Phe cac His gcg Ala tga g cc c tg	tat Tyr aat Asn gac Asp 1 aata acag acat acag	tac Tyr ttg Leu -15 ccg Pro tgg tggaa tcct	agt Ser -30 ggc Gly tct Ser gtta cac aaa rac	Met -45 ttt Phe gag Glu ggg Gly aagt acag acag atcag	Asr ac to a cara a	ser tttt Phe ttc Phe ttc Thr ctga aaga gcat gcttg	agg Arg ctt Leu -10 ttc Phe gcagc	10: 15 19 25 31 37 43

```
<210> 274
<211> 455
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 115..231
<221> sig_peptide
<222> 115..180
<223> Von Heijne matrix
      score 5
      seq HLFVTWSSQRALS/HP
<221> polyA_signal
<222> 419..424
<221> polyA site
<222> 445..455
<400> 274
aacctgccag tkatgcaaat gccaaaatgt gggtcatcat atagtatatt tgaaaccttt
                                                                       60
ctgaacatgt acaccacca atgctagagg ctgacttgga aaccggtggg tgca atg
                                                                      117
ccc gag gct gtg gaa caa tca gcc cat ctc ttt gtg acc tgg agc agt
                                                                      165
Pro Glu Ala Val Glu Gln Ser Ala His Leu Phe Val Thr Trp Ser Ser
                                             -10
                         -15
    -20
cag agg gcc ctc agt cac ccc gcc cca ttc ctc acc ara raa aar aat
                                                                      213
Gln Arg Ala Leu Ser His Pro Ala Pro Phe Leu Thr Xaa Xaa Lys Asn
                                    5
                                                                      261
cca ttt cta tgg aag ctc tgacgtaact tcagtgtttt ctacaatact
Pro Phe Leu Trp Lys Leu
            15
cctcctgccc cgccccatta aaacagttct tttgttaaaa aatavcctaa tggtccaact
                                                                      321
ttgctgtctg ttcttccaaa tgtttataat acacattatt tataaatatg tctgtttggg
                                                                      381
aagctaagaa caagctagtt tttacaacac aaatggaaat aaatgcaatt attataaaaa
                                                                      441
                                                                      455
tycaaaaaaa aaaa
```

<222> 662..673

<400> 275	60
attragetty cagactgeet tetateccag aacagetgag aaatetatga agetgagatt	120
ctgaaggacc cagettaggt tettecaett aggeeteaat teeetteett ttecagggge	180
agestagtt teccatgges etgaaacaca cacatttees estreette ecagaagesa	237
ctggccccc atagcaccca gtgcatcctt tttacaagtg gaagaactag g atg gct  Met Ala	
	285
ttc caa agt ctt cta gaa atg aag ttc ttt ctc tgt gca gct ttc ccc	
Phe Gln Ser Leu Leu Glu Met Lys Phe Phe Leu Cys Ala Ala Phe Pro	
-20 -15 -10	333
ctt gga gca gga gtg aag atg ttt cat tat ctt ggg cct ggg aaa cca	333
Leu Gly Ala Gly Val Lys Met Phe His Tyr Leu Gly Pro Gly Lys Flo	
_ e 1 5 ±0	207
att day dag get tet eee tee eee cae eee cat agg ame agg att tgg	381
Leu Xaa Gln Ala Ser Pro Ser Pro His Pro His Arg Xaa Arg Ile Trp	
15 20 25	
cet tagettetgg geetatesge tgeetteect ettytteeta ecaectette	434
Pro tgccttcctt trawctctgt tgggcttggg gatcttagtt ttcttttgtt tatttcccat	494
ctcatttttt tottotggto agtttttta agggggggtg ttgtggtttt ttgtttttgt	554
tttgcttctg aaaarcatt tgcctttctt cctctcccaa cataacaatc gtggtaacag	614
tttgcttctg aaaaarcatt tgcctttctt ctttcttaa aaaagaaaaa	673
aatgegactg ctgatttace gatgtattta atgtaagtaa aaaaaggaaa aaaaraaaa	
·	
<210> 276	
<211> 639	
<212> DNA	
<213> Homo sapiens	
2210 10110 000 101111	
<220>	
<221> CDS	
<222> 143427	
<221> sig_peptide	
<222> 143286	
<222> 143286 <223> Von Heijne matrix	
<222> 143286 <223> Von Heijne matrix score 7.5	
<222> 143286 <223> Von Heijne matrix	
<222> 143286 <223> Von Heijne matrix score 7.5	
<222> 143286  <223> Von Heijne matrix  score 7.5  seq FVILLLFIFTVVS/LV	
<222> 143286 <223> Von Heijne matrix score 7.5	
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix     score 7.5     seq FVILLLFIFTVVS/LV &lt;221&gt; polyA_signal</pre>	
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	60
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	60 120
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220 268
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220 268
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220 268 316
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220 268
<pre>&lt;222&gt; 143286 &lt;223&gt; Von Heijne matrix</pre>	120 172 220 268 316

15 20 25	
agt gaa ccc aac cct ctt ara akt atg atg gac aac atc aga aaa cgt	412
Ser Glu Pro Asn Pro Leu Xaa Xaa Met Met Asp Asn Ile Arg Lys Arg	
30 33	467
gaa act gaa gtg gtc taacactcta taraaaatga acaaaatctc tgaaagcagc	
Glu Thr Glu Val Val	
45	527
tcaacctctt ctgaraaaaa aaatatattc tgaggccaac tgttgctaca aaacaaattc	587
tgactgaatg gttaaaacat ttctagtara aggggaaaaa aaakttaaac atgcactgtt	639
tgtgtgtata sccatttcat taaatataca gtaaaactyc aaaaaaaaaa aa	033
<210> 277	
<211> 772	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 284463	
<221> sig_peptide	
<222> 284379	10
<223> Von Heijne matrix	
score 3.79999995231628	
seq TFINITLWLGSLC/QR	4
<221> polyA_site	
<222> 762772	
<400> 277	60
acagetgggg ctttgtette tttattgeta ggagaatgta gcaatagaag tteteatege	120
cotgtattgc actittggtt traaggactg gacccagagt tootgaaagc caaactccat	180
aagetgetea gtaagtteea ageacatage eggetkhggg atgegatteg gtegaggtet	240
gttgaatgaa ggtagacgca gcaggcagtt tgtccttacc agtgacctgg aagacggtgg	295
cactteetga gtgageteae ttacetteee tgaatggtga gge atg gat gaa tat  Met Asp Glu Tyr	2,55
-30	
	343
tee tgg tgg tge cae gtg tta gag gtg gta aag ggt caa atg ttt act	243
Ser Trp Trp Cys His Val Leu Glu Val Val Lys Gly Gln Met Phe Thr	
-25 -20 -15	391
ttt att aat att aca tta tgg ctt ggt tct ctg tgt cag cga ttt ttc	371
Phe Ile Asn Ile Thr Leu Trp Leu Gly Ser Leu Cys Gln Arg Phe Phe	
-10 -5	439
tat gcc tcg ggt act tat ttc cta ata tat atc agc aca gta acg cct	433
Tyr Ala Ser Gly Thr Tyr Phe Leu Ile Tyr Ile Ser Thr Val Thr Pro	
5 10 15 20	403
agc tgg agg ctt tgt ctt gtt agt tgataaatta gtggtaacag gtagatttgg	493
Ser Trp Arg Leu Cys Leu Val Ser	
25	
ttacctccca aagtgctggg attrcagacg tgagccaccg cgcctggccg aaacaattct	553
tttgaaagag agaagtetee etgtgttgeg eaggetggte teagacteet ggggteaagt	613
gageeteetg etttegeete etaaagtget gggattacag gegtgageea eegeaceegg	673
acagatgtgt tgattttaaa gtgggtatga ggcctgagcc ctggagtttg agaccagcct	733
ggacaacatg gcaagaccct gtctctccaa aaaaaaaaa	772

<210> 278 <211> 840 <212> DNA

<213> Homo sapiens																
<220: <221: <222:	> CDS		71													·
<pre>&lt;221&gt; sig_peptide &lt;222&gt; 162398 &lt;223&gt; Von Heijne matrix</pre>																
<221> polyA_signal																
<221 <222	-	-														
<400	> 27	8								~~t	+ < > <	cata	מכ מ	agat	adcad	60
aaaa	actg	ag g	cctg	ggag	c ag	gaac	teta	agg	ggaa	caa	aaat	caca	gg a	tgto	agcag	120
atgt	aacg ttcc	cg g	gaag	gagg	a aa c gg	ataa	aggg	gtc	ccag	cac	c at	g ga	.g ga		g aac o Asn -75	176
cct	gaa	aaa	aac	atg	aaq	caq	cag	gat	tca	ccc	aag	gag	aga	agt	ccc	224
Pro	Glu	Glu	Asn	Met -70	Lys	Gln	Gln	Asp	Ser -65	Pro	ьуs	GIU	Arg	-60	PIO	
cag	agc	cca	gga	ggc	aac	atc	tgc	cac	ctg	999	gcc	ccg	aag	tgc	acc	272
Gln	Ser	Pro	Gly -55	Gly	Asn	Ile	Cys	His -50	Leu	GIY	Ala	Pro	-45	Cys	1111	320
Arg	Cys	Leu	Ile	acc Thr	Phe	Ala	Asp -35	Ser	Lys	Phe	GIN	-30	Arg	HIS	Mec	
aag Lys	Arg	a a a	cac His	cca Pro	gcg Ala	gac Asp -20	ttc Phe	gtg Val	gcc Ala	cag Gln	aag Lys -15	ctg Leu	cag Gln	gjà aaa	gtc Val	368
ctc	-25 ttc	atc	tac	ttc	acc	tac	acc	cqc	tcc	ttc	ccc	tcc	tcc	aaa	gcc	416
Leu -10	Phe	Ile	Cys	Phe	Thr	Cys	Ala	Arg	Ser	Phe 1	Pro	Ser	Ser	Lys 5	Ala	
ckr	rkc	acc	cac	car	cgc	agc	cac	ggt	cca	rcc	gcc	aag	CCC	acc	ctg	464
Xaa	Xaa	Thr	His	Gln	Arg	Ser	His	Gly 15	Pro	Xaa	Ala	гуs	20	THE	Leu	
ccg	gtt	gca	acc	act	act	gcc	car	CCC	acc	ttc	cct	tgt	cct	gac	tgt	512
Pro	Val	Ala 25	Thr	Thr	Thr	Ala	Gln 30	Pro	Thr	Phe	Pro	Cys 35	Pro	Asp	Cys	560
ggc	Lys	acc Thr	ttt Phe	GJA 333	cag Gln	gct Ala 45	gtt Val	tct Ser	ctg Leu	arg Xaa	cgg Arg 50	cac His	csc Xaa	Gln	Xaa	300
cat	40	atc	cat	gcc	cct	cct	aac	acc	ttc	gcc	tgc	aca	rad	tgc	ggt	608
His	Glu	Val	Arg	Ala	Pro 60	Pro	Gly	Thr	Phe	Ala 65	Cys	Tnr	хаа	Cys	70	
cag	gac Asp	ttt Phe	gct Ala	car Gln	gaa Glu	rca Xaa	Gly 999	ctg Leu	cat His 80	caa Gln	cac His	tac Tyr	att Ile	cgg Arg 85	cat His	656
			Gly	75 ctc Leu	tga	gttc	agc	ttaa		ct c	cacg	gtga	c gg		ctct	711
<b>~</b> + ~	aa+~	~+ ·	90	+ 0 = 0	CC 2	tast	atoo	a at	acsa	gaac	tct	agaa	qcc	ctga	aggatt	771
grg tac	ttee	gta ctc	ccct	ggga	ag g	caga	gggc	t ct	taat	aaag	agg	accc	aka	agat	tcttaa	831
	aaaa		•	J		-										840

```
<210> 279
<211> 840
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 63..632
<221> sig_peptide
<222> 63..308
<223> Von Heijne matrix
      score 4.40000009536743
      seg NLPHLQVVGLTWG/HI
<221> polyA_signal
<222> 808..813
<221> polyA_site
<222> 829..840
<400> 279
aactteeggt egegeeaseg ecegttgeea gttetgegeg tgteetgeat etceagtatg
                                                                       60
ga atg tat gtd tgg ccc tgt gct gtg gtc ctg gcc cag tac ctt tgg
                                                                      107
   Met Tyr Val Trp Pro Cys Ala Val Val Leu Ala Gln Tyr Leu Trp
                               -75
ttt cac aga aga tct ctg cca ggc aag gcc atc tta gag att gga gct
                                                                       155
Phe His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala
                                               -55
                             -60
        -65
gga gtg agc ctt cca gga att ttg gct gcc aaa tgt ggt gca gaa gta
                                                                       203
Gly Val Ser Leu Pro Gly Ile Leu Ala Ala Lys Cys Gly Ala Glu Val
                                             -40
                        -45
ata ctg tca gac agc tca gaa ctg cct cac tgt ctg gaa gtc tgt cgg
                                                                       251
Ile Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg
                                         -25
                     -30
-35
caa ago tgo caa atg aat aac otg coa cat otg cag gtg gta gga cta
                                                                       299
Gln Ser Cys Gln Met Asn Asn Leu Pro His Leu Gln Val Val Gly Leu
                                     -10
                -15
                                                                       347
aca tgg ggt cat ata tct tgg gat ctt ctg gct cta cca cca caa gat
Thr Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp
                            - 5
 att atc ctt gca tct gat gtg ttc ttt gaa cca gaa rat ttt gaa gac
 Ile Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Xaa Phe Glu Asp
                                             25
                        20
 att ttg gct aca ata tat ttt ttg atg cac aar aat ccc aag gtc caa
 Ile Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln
                                         40
                     35
 ttg tgg tct act tat caa gtt agg art gct gac tgg tca ctt gaa gct
                                                                       491
 Leu Trp Ser Thr Tyr Gln Val Arg Xaa Ala Asp Trp Ser Leu Glu Ala
                                     55
                50
 tta ctc tac aaa tgg gat atg aaa tgt gtc cac att cct ctt gag tct
                                                                       539
 Leu Leu Tyr Lys Trp Asp Met Lys Cys Val His Ile Pro Leu Glu Ser
                                 70
             65
 ttt gat gca gac aaa gaa rat ata gca gaa tct acc ctt cca gga aga
                                                                       587
 Phe Asp Ala Asp Lys Glu Xaa Ile Ala Glu Ser Thr Leu Pro Gly Arg
                             85
 cat aca gtt gaa atg ctg gtc att tcc ttt gca aag gac agt ctc
                                                                       632
 His Thr Val Glu Met Leu Val Ile Ser Phe Ala Lys Asp Ser Leu
                                             105
                         100
 tgaattatac ctacaacctg ttctgggaca gtatcaatac tgatgagcaa cctggcacac
                                                                       692
 aaactatgag cagaccactt cagcttgaga atgcagtggg tctgaagatg gtcaagtctg
                                                                       752
```

tttg: aata	cctta cgtat	ar a t a	tttt	gatgt caaaa	ca a aa	ccta aaaa	gaca aa	aca	ctta	aac	tcat	atga	aa c	aaaa	attaa	812 840 .
<211 <212	> 280 > 849 > DNA > Hor	9 <b>A</b> .	apie	ns												
	> > CDS > 21		2													
<222	sc	20 n He ore	0 ijne 4.80	e mat: 0000: KSQT	1907											
<222	> po > 82	18	26													
	> po > 83		-	<b>!</b>												
<400 agta	> 28 agtc	0 CC C	ecgo	ctcg	c at Me	t Me	g go et Al	t go .a Al	eg gt La Va	ig co al Pi -5	o Pr	g gg o Gl	c ct y Le	g ga eu Gl	g ccg u Pro -50	53
tgg Trp	aac Asn	cgt Arg	gtg Val	aga Arg -45	atc	cct	aag Lys	gcg Ala	999 Gly -40	aac	cgc	agc Ser	gca Ala	gtg Val -35	aca Thr	101
gtg Val	cag Gln	aac Asn	ccc Pro -30	ggc Gly	gcg Ala	gcc Ala	ctt Leu	gac Asp -25	ctt Leu	tgc Cys	att Ile	gca Ala	gct Ala -20	gta Val	att Ile	149
Lys	Glu	Cys -15	His	ctc Leu	Val	Ile	Leu -10	Ser	Leu	Lys	Ser	Gln -5	Thr	Leu	Asp	197
gca Ala	gaa Glu l	aca Thr	gat Asp	gtg Val	tta Leu 5	tgt Cys	gca Ala	gtc Val	ctt Leu	tac Tyr 10	agc Ser	aat Asn	cac His	aac Asn	aga Arg 15	245
Met	Gly	Arg	His	aaa Lys 20	Pro	His	Leu	Ala	Leu 25	Lys	Gln	Val	Glu	30	Cys	293
Leu	Lys	Arg	Leu 35	aaa Lys	Asn	Met	Asn	Leu 40	Glu	Gly	Ser	Ile	Gln 45	Asp	Leu	341
Phe	Glu	Leu 50	Phe	tct Ser	Ser	Lys										392
gaat ktgt cate	tataa tggta	acc a at a	aatta cttc: tata:	ataco tgaca ttaaa	w call	agct; gatc; atca;	gtak: tatg: gtgg:	a aa g ga: g ct:	twtt rtga gtta	gttt ctgg ttgt	taa: tgt: gct:	tgtgg gacat taact	ggg ttg tac	tacc; aaat ctca	ckgttg yggtgt ctgggt agttga atgaaa	452 512 572 632 692
gta	atggg gtgct	gar :	mttg ttct	cacat	a g	ctga cttg	aaat ccct	g tg a tg	aagg catc	gtcg tctt	CCC	aggga	agg	amat	ggaagc	752 812 849

```
<210> 281
<211> 1344
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 21..503
<221> sig_peptide
<222> 21..344
<223> Von Heijne matrix
     score 5.30000019073486
     seq ACMTLTASPGVFP/SL
<221> polyA_signal
<222> 1305..1310
<221> polyA site
<222> 1330..1341
<400> 281
                                                                      53
aaacaactcc ggaaagtaca atg acc agc ggg cag gcc cga gct tcc wyc cag
                      Met Thr Ser Gly Gln Ala Arg Ala Ser Xaa Gln
                                  -105
tcc ccc cag gcc ctg gag gac tcg ggc ccg gtg aat atc tca gtc tca
                                                                      101
Ser Pro Gln Ala Leu Glu Asp Ser Gly Pro Val Asn Ile Ser Val Ser
                           -90
                                         -85
        -95
ate ace eta ace etg gac eca etg aaa ece tte gga ggg tat tee ege
Ile Thr Leu Thr Leu Asp Pro Leu Lys Pro Phe Gly Gly Tyr Ser Arg
                                            -70
                        -75
                                                                      197
aac gtc acc cat ctg tac tca acc atc tta ggg cat cag att gga ctt
Asn Val Thr His Leu Tyr Ser Thr Ile Leu Gly His Gln Ile Gly Leu
-65
                                        -55
                    -60
tca ggc agg gaa gcc cac gag gag ata aac atc acc ttc acc ctg cct
                                                                      245
Ser Gly Arg Glu Ala His Glu Glu Ile Asn Ile Thr Phe Thr Leu Pro
                                                                      293
aca gog tgg ago tca gat gac tgc gcc ctc cac ggt cac tgt gag cag
Thr Ala Trp Ser Ser Asp Asp Cys Ala Leu His Gly His Cys Glu Gln
                                -25
                                                    -20
                                                                      341
gtg gta ttc aca gcc tgc atg acc ctc acg gcc agc cct ggg gtg ttc
Val Val Phe Thr Ala Cys Met Thr Leu Thr Ala Ser Pro Gly Val Phe
                            -10
                                                                      389
ccg tca ctg tac agc cac cgc act gtg ttc ctg aca cgt aca gca acg
Pro Ser Leu Tyr Ser His Arg Thr Val Phe Leu Thr Arg Thr Ala Thr
                                        10
                                                                      437
cca cgc tct ggt aca aga tct tca caa ctg cca gag atg cca aca caa
Pro Arg Ser Gly Thr Arg Ser Ser Gln Leu Pro Glu Met Pro Thr Gln
                20
                                    25
                                                                      485
aat acg ccc aaa att aca atc ctt tct ggt gtt ata agg ggg cca ttg
Asn Thr Pro Lys Ile Thr Ile Leu Ser Gly Val Ile Arg Gly Pro Leu
                                40
            35
                                                                      533
gaa aag tot ato atg oft taaatcocaa gottacagtg attgttccag
Glu Lys Ser Ile Met Leu
atgatgaccg ttcattaata aatttgcatc tcatgcacac cagttacttc ctctttgtga
tggtgataac aatgttttgc tatgctgtta tcaagggcag acctagcaaa ttgcgtcaga
                                                                      713
gcaatcctga attttgtccc gagaaggtgg ctttggctga agcctaattc cacagctcct
tgttttttga gagagactga gagaaccata atccttgcct gctgaaccca gcctgggcct
                                                                      773
ggatgctctg tgaatacatt atcttgcgat gttgggttat tccagccaaa gacatttcaa
                                                                     833
                                                                      893
gtgcctgtaa ctgatttgta catatttata aaaatctatt cagaaattgg tccaataatg
cacgtgcttt gccctgggta cagccagagc ccttcaaccc caccttggac ttgaggacct
```

acctgatggg acgtttccac gtgtctctag agaaggatcc tggatctagc tggtcacgac

acctgatggg acgtttccac gtgtctctag agaaggatcc tggatctagc tggtcacgac gatgttttca ccaaggtcac aggagcattg cgtcgctgat ggggttgaag tttggtttgg	1013 1073 1133 1193 1253 1313 1344
<210> 282 <211> 671 <212> DNA <213> Homo sapiens	
<220> <221> CDS . <222> 1201	
<pre>&lt;221&gt; sig_peptide &lt;222&gt; 163 &lt;223&gt; Von Heijne matrix</pre>	
<221> polyA_signal <222> 637642	
<221> polyA_site <222> 660671	
<pre>&lt;400&gt; 282 atg ctg gga ggt gac cat agg gct ctg ctt tta aag ata tgg ctg ctt Met Leu Gly Gly Asp His Arg Ala Leu Leu Leu Lys Ile Trp Leu Leu -20 -15 -10</pre>	48
caa agg cca gag tca cag gaa gga ctt ctt cca ggg aga tta gtg gtg Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro Gly Arg Leu Val Val -5 1 5 10	96
atg gag agg aga gtt aaa aat gac ctc atg tcc ttc ttg tcc acg gtt Met Glu Arg Arg Val Lys Asn Asp Leu Met Ser Phe Leu Ser Thr Val 15 20 25	144
ttg ttg agt ttt cac tct tct aat gca agg gtc tca cac tgt gaa cca Leu Leu Ser Phe His Ser Ser Asn Ala Arg Val Ser His Cys Glu Pro 30 35 40	192
ctt agg atg tgatcacttt caggtggcca ggaatgttga atgtctttgg Leu Arg Met 45	241
ctcagttcat ttaaaaaaga tatctatttg aaagttctca rarttgtaca tatgtttcac	301
agtacaggat ctgtacataa aagtttcttt cctaaaccat tcaccaagag ccaatatcta	361
ggcattttct tggtagcaca aattttctta ttgcttaraa aattgtcctc cttgttattt	421 481
ctgtttgtaa racttaagtg agttaggtct ttaaggaaag caacgctcct ctgaaatgct tgtctttttt ctgttgccga aatarctggt cctttttcgg gagttaratg tatarartgt	541
ttgtatgtaa acatttcttg taggcatcac catgaacaaa gatatattt ctatttattt	601
attatatgtg cacttcaaga agtcactgtc agagaaataa agaattgtct taaatgtcaa aaaaaaaaa	661 671

<210> 283

<211> 1601

<212> DNA

<213> Homo sapiens

<220> <221> CDS <222> 39..1034 <221> sig_peptide <222> 39..134 <223> Von Heijne matrix score 6.09999990463257 seq LPLLTSALHGLQQ/QH <221> polyA_signal <222> 1566..1571 <221> polyA_site <222> 1587..1597 <400> 283 agccccagat cctgaaggag gtgcagagcc cagagggg atg atc kcg ctg agg gac 56 Met Ile Xaa Leu Arg Asp 104 aca gct gcc tcc ctc cgc ctt gag aga gac aca agg cag ttg cca ctg Thr Ala Ala Ser Leu Arg Leu Glu Arg Asp Thr Arg Gln Leu Pro Leu -20 -15 152 ctc acc agt gcc ctg cac gga ctg cag cag cac cca gcc ttc tct Leu Thr Ser Ala Leu His Gly Leu Gln Gln His Pro Ala Phe Ser - 5 ggt gtg gca cgg ctg gcc aag cgg tgg gtg cgt gcc cag ctt ctt ggt 200 Gly Val Ala Arg Leu Ala Lys Arg Trp Val Arg Ala Gln Leu Leu Gly 15 10 gag ggt ttc gct gat gag agc ctg gat ctg gtg gcc gct gcc ctt ttc 248 Glu Gly Phe Ala Asp Glu Ser Leu Asp Leu Val Ala Ala Ala Leu Phe 30 ctg cac cct gag ccc ttc acc cct ccg agt tcc ccc cag gtt ggc ttc 296 Leu His Pro Glu Pro Phe Thr Pro Pro Ser Ser Pro Gln Val Gly Phe 45 ctt cga ttc ctt ttc ttg gta tca acg ttt gat tgg aag aac aac ccc 344 Leu Arg Phe Leu Phe Leu Val Ser Thr Phe Asp Trp Lys Asn Asn Pro 60 392 ctc ttt gtc aac ctc aat aat gag ctc act gtg gag gag cag gtg gar Leu Phe Val Asn Leu Asn Asn Glu Leu Thr Val Glu Glu Gln Val Glu 80 440 ate ege agt gge tte etg gea get egg gea eag ete eee gte atg gte Ile Arg Ser Gly Phe Leu Ala Ala Arg Ala Gln Leu Pro Val Met Val 95 488 att gtt acc ccc caa rac cgc aaa aac tct gtg tgg aca cag gat gga Ile Val Thr Pro Gln Xaa Arg Lys Asn Ser Val Trp Thr Gln Asp Gly 110 536 ccc tca gcc car atc ctg cag cag ctt gtg gtc ctg gca gct gaa scc Pro Ser Ala Gln Ile Leu Gln Gln Leu Val Val Leu Ala Ala Glu Xaa 125 ctq ccc atq tta rar aas cag ctc atg gat ccc cgg gga cct ggg gac 584 Leu Pro Met Leu Xaa Xaa Gln Leu Met Asp Pro Arg Gly Pro Gly Asp 145 140 atc agg aca gkg ttc cgg ccg ccc ttg gac att tac gac gtg ctg att 632 Ile Arg Thr Xaa Phe Arg Pro Pro Leu Asp Ile Tyr Asp Val Leu Ile 160 155 cgc ctg tct cct cgc cat atc ccg cgg cac cgc cag gct gtg gac tcr 680 Arg Leu Ser Pro Arg His Ile Pro Arg His Arg Gln Ala Val Asp Ser 175 170

cca gct gcc tcc ttc tgc cgg ggc ctg ctc agc cag ccg ggg ccc tca

Pro Ala Ala Ser Phe Cys Arg Gly Leu Leu Ser Gln Pro Gly Pro Ser

185		190		195	
	ccc ata cta		gat cct cct	cag ctc tat	ctg acg 776
				Gln Leu Tyr	
200		205		210	
				ttc ttc tat	
		Gly Asp L		Phe Phe Tyr	
215	220		225		230
				ccc acc agc	
HIS GIA GIA		GIY VAI L	Leu Trp Lys 240	Pro Thr Ser	245
CCG C2G CCG	235	tec age a		cgc atg gtg	
				Arg Met Val	
110 0111 110	250		255	260	
cga ggt ggg		atg gtg c	cc aat gtt	gaa gca atc	ctg gag 968
				Glu Ala Ile	
265		270		275	
				act gtg gag	
_	Val Leu Gly		Leu Val Gln	Thr Val Glu	Ala Arg
280		285		290	2004
		-	gc tctggagca	a gctgtagacg	1064
Ser Giu Arg	Trp Thr Val				
	_		atgtcagtag	gatgacetee a	ccctccttq 1124
				tgaatcatct c	
				ggggcatggt g	
				agctcagaga a	
				tccctgcgat t	
tttcttagtc t	ttcttccaga a	acagagaag	gggatgtgtg	cctgggagag g	
		_		ttcactgcag t	
				atcaccgaca c	
gaaaggggct a	atatgtgtat g	aatagacca	cattgaagga	gcaaaaaaaa a	aamcch 1601

<211> 1206

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 69..263

<221> sig_peptide

<222> 69..125

<223> Von Heijne matrix score 3.90000009536743 seq ALSMSSFSFHSSS/CS

<221> polyA_signal

<222> 1173..1178

<221> polyA_site

<222> 1196..1205

<400> 284

60 acatttgtga ctttaccaat acceteccag ttettgatag acagetgtag gttgetgggt tcaagaat atg ggt ggg ata tgg aat gct ctt tca atg tct agc ttc agt 110 Met Gly Gly Ile Trp Asn Ala Leu Ser Met Ser Ser Phe Ser -15 158

ttt cat tca tcc tcc tgc tca gca ctg tca gcc aag agc tta ctc agc Phe His Ser Ser Ser Cys Ser Ala Leu Ser Ala Lys Ser Leu Leu Ser

-5 1 5 10	
aga cac cac ata ctg cag cag ttc cta gtg aga aaa tct gtg cca cta	206
Arg His His Ile Leu Gln Gln Phe Leu Val Arg Lys Ser Val Pro Leu	
15 20 25	
gaa aat gct tca ctt cca ttt cct cac ctg ggc agt tct ctg ttt aaa	254
Glu Asn Ala Ser Leu Pro Phe Pro His Leu Gly Ser Ser Leu Phe Lys	
30 35 40	
att gtg ggc tgatttggtc ttcctctcct cctcccactg ttactgccct.	303
Ile Val Gly	
45	
gcagcccttg ttcaggtgta cagaccctta ttctggcctc tagtgtcctt gtctgtcatg	363
acacaccett cegeceaaat acetetgace ecaaggetgg aatggggetg gtaggarata	423
agtttgctta ctcatartca tgtcctttct cttggcacct gcttccctgc ggtgtcctca	483
aatggatttc tgtgtggcag tggartgatt gcatgaattt ttctgtaaca cattaacttt	543
gtattattat taagggartt tgaraaagct ttgcttataa tgtcaaggca aggaggtaaa	603
aactggagcc caaakaaatt cccttagggc aagattatgt tataataraa aattgaattt	663
cctgaggcag tggctgccac cccttttcar atgtttagtc ctgcaaatag catctttctt	723
gtagtctgtg acatggatgg ggatgctagg gcccttaggg gcaaggggac taaactaaat	783
caakttgagt ttttttccag caggggttar gggaggtact csctgttgat atttgacact	843
araaagtaat cttttttaca aaactgtttt tctaggtggg tggaaagtga aactgccaca	903
tccttgttgg tttagtccaa raratcattt gcaacaacag taratgtccg ggttttgttt	963
ctgtcttttt attatgaaaa actatgttaa gggggaaaat gtggattatg gtaaccarag	1023
gaatccctas cettgtttte ettaraarae ttgtttagtg ttttatcara egtetgttgt	1083
agttgtarac aggaaagctt gtgaraaaaa caccacatgg ascctgtaaa tgtttttgca	1143
caacctgtaa agcattettg gaaktggeea gtaaaaaggg gttttaccat ttaaaaaaaa	1203
aat	1206
<210> 285	
<211> 536	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 115285	
<221> sig_peptide	
<222> 115204	
<223> Von Heijne matrix	
score 3.7000004768372	
seq SMMLLTVYGGYLC/SV	
221 males aireas	
<221> polyA_signal	
<222> 505510	
1993, males aire	
<221> polyA_site <222> 525536	
<2227 323336	
<400> 285	
acgagtgctg cgttcggctg tgctgggaag ttgcgtagac agtggcctcg agaccctgcc	60
tgcctgagga ggcctcggtt ggatgcgaag gagctgcagc atccagggga caag atg	117
Met	
-30	
cca act ggc aag cag cta gct gac att ggc tat aag acc ttc tct acc	165
Pro Thr Gly Lys Gln Leu Ala Asp Ile Gly Tyr Lys Thr Phe Ser Thr -25 -20 -15	
	212
Ser Met Met Leu Ter Val Ter Cly Cly Ter Leu Ger Ser Val Des	213
Ser Met Met Leu Thr Val Tyr Gly Gly Tyr Leu Cys Ser Val Arg	
- The state of the	261
gtc tac cac tat ttc cag tgg cgc agg gcc cag cgc cag gcc gca gaa	261

Val Tyr His Tyr Phe Gln Trp Arg Arg Ala Gln 5	Arg Gln Ala Ala Glu 15
gaa cag aag dac tca gga atc atg tagaactggg Glu Gln Lys Xaa Ser Gly Ile Met 20 25	-
asakgcccaa ggcatgctgt ggagagactt cacctgccac	catttccagg tcaacaggac 375
tagagegttg atggttttca aaccetgttg gaagaaagtg	cccatggttt ctctggttct 435
gccartttga cagtttatgg argcttttga atcgtaatar	
cctacagaca ttaaataatt tgctgtgtca aaaaaaaaaa	a 536
<210> 286	
<211> 529	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 90344	
<221> sig_peptide	
<222> 90140 <223> Von Heijne matrix	
score 8.19999980926514	
seq LLLITAILAVAVG/FP	
•	
<221> polyA_signal	
<222> 500505	•
<221> polyA_site	
<222> 515527	
<400> 286	
aatatrarac agctacaata ttccagggcc artcacttgc	catttctcat aacagcgtca 60
gagagaaaga actgactgar acgtttgag atg aag aaa	gtt ctc ctc ctg atc 11
Met Lys Lys -15	Val Leu Leu Ile -10
aca gcc atc ttg gca gtg gct gtw ggt ttc cca	
Thr Ala Ile Leu Ala Val Ala Val Gly Phe Pro	Val Ser Gln Asp Gln
-5	5
gaa cga gaa aaa aga agt atc agt gac agc gat	gaa tta gct tca ggr 209
Glu Arg Glu Lys Arg Ser Ile Ser Asp Ser Asp	Glu Leu Ala Ser Gly
10 15	- 20
wtt ttt gtg ttc cct tac cca tat cca ttt cgc	
Xaa Phe Val Phe Pro Tyr Pro Tyr Pro Phe Arg	
25 30 cca ttt cca aga ttt cca tgg ttt aga cgt aat	35 ttt cct att cca ata 309
Pro Phe Pro Arg Phe Pro Trp Phe Arg Arg Asm	
40 45 50	55
cet gaa tet gee eet aca act eee ett eet age	gaa aag taaacaaraa 354
Pro Glu Ser Ala Pro Thr Thr Pro Leu Pro Ser	
60 65	
ggaaaagtca crataaacct ggtcacctga aattgaaatt	
caaaattcct gttaataaaa raaaaacaaa tgtaattgaa	
gtcaatatct ttagtgatct tctttaataa acatgaaagc	aaaaaaaaa aaacc 52

<210> 287 <211> 493

<212> DNA

PCT/IB98/02122 -

<213> Homo sapiens <220> <221> CDS <222> 57..311 <221> sig_peptide <222> 57..107 <223> Von Heijne matrix score 8.19999980926514 seq LLLITAILAVAVG/FP <221> polyA signal <222> 467..472 <221> polyA_site <222> 482..493 <400> 287 aacttgccat ttctcataac agcgtcagag agaaagaact gactgaaacg tttgag atg 59 Met 107 aag aaa gtt ctc ctc ctg atc aca gcc atc ttg gca gtg gct gtt ggt Lys Lys Val Leu Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val Gly -15 -10 - 5 ttc cca gtc tct caa gac cak gaa cga gaa aaa aga agt atc agt gac 155 Phe Pro Val Ser Gln Asp Xaa Glu Arg Glu Lys Arg Ser Ile Ser Asp 10 age gat gaa tta get tea ggg ttt ttt gtg tte eet tae eea tat eea 203 Ser Asp Glu Leu Ala Ser Gly Phe Phe Val Phe Pro Tyr Pro Tyr Pro 20 ttt cgc cca ctt cca cca att cca ttt cca aga ttt cca tgg ttt aga 251 Phe Arg Pro Leu Pro Pro Ile Pro Phe Pro Arg Phe Pro Trp Phe Arg 40 299 cgt aat ttt cct att cca ata cct gaa tct gcc cct aca act ccc ctt Arg Asn Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Thr Thr Pro Leu 55 60 ccg agc gaa aag taaacaagaa ggaaaagtca cgataaacct ggtcacctga 351 Pro Ser Glu Lys aattgaaatt gagccacttc cttgargaat caaaattcct gttaataaaa gaaaaacaaa 411 tgtaattgaa atagcacaca gcatteteta gteaatatet ttagtgatet tetttaataa 471 acatgaaagc aaaaaaaaa aa 493 <210> 288 <211> 521

<211> 521
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 96..302
<221> sig_peptide
<222> 96..182
<223> Von Heijne matrix
score 5
seq ELSLLPSSLWVLA/TS
<221> polyA_site

<222> 501..514

<400> 288	
aagagacgtc accggctgcg cccttcagta tcgcggacgg aaga	tageat coaccaccea 60
aagagacgee acceptings continues regard at a see t	-55-555
teteatecag eggetgegga aetgggegte eggge atg ace t	ys Arg Gly Ser
Met III C	-25
	_
tgc agc tac gct acc agg aga tct cca agc gaa ctc	age ete ete eta ioi
Cys Ser Tyr Ala Thr Arg Arg Ser Pro Ser Glu Leu	ser Leu Leu Pro
-20 -15	-10
ago too etg tgg gto eta geo aca ago tot eca aca	att act att gca 209
Ser Ser Leu Trp Val Leu Ala Thr Ser Ser Pro Thr	Ile Thr Ile Ala
-5 1 5	
ctc gcg atg gcc gcc ggg aat ctg tgc ccc ctt cca	tca tca tkt cgt 257
Leu Ala Met Ala Ala Gly Asn Leu Cys Pro Leu Pro	Ser Ser Xaa Arg
10 15 20	25
crc aaa agg cgc tgg tgt cag gca asc car caa ara	act cta cta 302
Xaa Lys Arg Arg Trp Cys Gln Ala Xaa Gln Gln Xaa	Ala Leu Leu
	40
tagetgecae tgaaaaraag geggtgaete eageteetee eata	cgacta ctttqcctcc 422
cctcggacca gccttacctg tgacactgca ccctcacggc cacc	cgacta ctttgcctcc 422
ttggatttcc tccagggaga atgtgaccta atttatgaca aata	cgtara gctcaggtat 482
cacttctagt tttactttaa aaaataaaaa aatagagac	521
•	
<210> 289	
<211> 811	
<212> DNA	
<213> Homo sapiens	
<220>	•
<221> CDS	
<222> 161526	
<221> sig_peptide	
<222> 161328	
<223> Von Heijne matrix	
score 4.19999980926514	
seg XSPLLTLALLGQC/SL	
sed varuningmade, on	
DD1. malan mita	
<221> polyA_site	
<222> 799811	
<400> 289	
aaaaaattgc agtgctgaag acactggacc cgcaaaaggc tgtc	cctccc aaacctggga 60
ttctgggctc actgagttca cctgcgagtc agccctacct gcac	tgctct ggtctagtac 120
aaacaggctg ctggcattga ggtctgctac aaaaanarta atg	gtc cca tgg ccc 175
	Val Pro Trp Pro
	-55
and the second s	
agg ggc aag gtg aaa act gct cct att ccc atc tct	agg ttt ccc tto
Arg Gly Lys Val Lys Thr Ala Pro Ile Pro Ile Ser	Alg Phe Plo Phe
-50 -45 -40	
ctc cct acc cac gac cca ccc acc cca gca cat tgg	tot coa goa tot 271
Leu Pro Thr His Asp Pro Pro Thr Pro Ala His Trp	Ser Pro Ala Ser
-35 -30 -25	-20
cat cag cag ttt aaa cat kkg tca ccc ctc ctc act	ttg gcc ctg ctg 319
His Gln Gln Phe Lys His Xaa Ser Pro Leu Leu Thr	Leu Ala Leu Leu
-15 -10	<del>-</del> 5
ggt cag tgc tct ctg ttc arc aat ttg agg aaa aaa	ctt qca ggg caa 367
Gly Gln Cys Ser Leu Phe Xaa Asn Leu Arg Lys	Leu Ala Gly Gln
1 5	10
ass one ass ass the out too the too age of occ	

ھے

Lys Ala Lys Lys Leu Pro Ser Phe Ser Ser Leu Pro Leu Thr Leu Trp	
cca tta act cct caa ttt gct gag ctc act aca gtg gca caa aaa aaa Pro Leu Thr Pro Gln Phe Ala Glu Leu Thr Thr Val Ala Gln Lys Lys 30 35 40 45	463
ttg agg tgg tcc ggg acc cta ggt tgg ggt cca gtt ccc agc tgg gtt Leu Arg Trp Ser Gly Thr Leu Gly Trp Gly Pro Val Pro Ser Trp Val	511
caa ttt ttt tta ggg tgaatggagg garagttggg gactgaaaas ccttcaaara Gln Phe Phe Leu Gly 65	566
caatgttatt acagcaktct ccccttatcc aaaktttcct tttcctgadt ttcagttagc tatggtcaac cgcttggaaa atakttgaac acagtacaat aaratatttt gaggctggga ktggtggctc atgcctgtaa taatcccagg actttgtgar accaaktttg aaggatcact tgaacccagg aktttgarac cascctgggc aacatrgtra gacctcatct ctacaaaaaa aaaaa	626 686 746 806 811
<210> 290 <211> 625 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 210332	
<221> sig_peptide <222> 210299 <223> Von Heijne matrix score 8.10000038146973 seq ITCLLAFWVPASC/IQ	
<221> polyA_signal <222> 594599	
<221> polyA_site <222> 613625	
<pre>&lt;400&gt; 290 acaggtcsmc ttaacatctc ttgatttgag ccactcccac tgtcatcagc tttcacctgg attatcgtga cagcctccta ctgcttctct atcatgtggc cagagctatc ttccctaaaa atgcattgca tagttgatca agtcactctc tggcctaaaa ccttccttgg ctccctgctg ccctcaggat aaagtctgga cccctcagc atg gct tgt gag act cat ggt gtc</pre>	60 120 180 233
ctt gtc cct gct cac ctc tct ggt ctc atc act tgc ctt ctt gca ttc Leu Val Pro Ala His Leu Ser Gly Leu Ile Thr Cys Leu Leu Ala Phe -20 -15 -10	281
tgg gtc cca gcc tcc tgt atc cag aga tgc agt ggc tct cca ttg cca Trp Val Pro Ala Ser Cys Ile Gln Arg Cys Ser Gly Ser Pro Leu Pro -5 1 5 10	329
ctc tgattcctcc tttcttttgg tcacagagaa agggtacttt ctctgtcaaa Leu	382
teteaactta gaettgaett eeteeaagga getttggeta taetetetee ewegaeeeee aeeetggeat aetacaeara teaetetggg eteaettgee tgeetaatgg teateteeee agtaaactgt aageteettg agggeaagga ttgtgttgga atttttgtat taacagtgee tggettggtg cetggeaeet aaaaageaet caataaatgt ttgtttaatg aaaaaaaaaa aaa	442 502 562 622 625

```
<210> 291
 <211> 684
 <212> DNA
 <213> Homo sapiens
 <220>
 <221> CDS
 <222> 212..361
 <221> sig_peptide
 <222> 212..319
 <223> Von Heijne matrix
       score 4.09999990463257
      seq HWLFLASLSGIKT/YQ
 <221> polyA_signal
 <222> 650..655
<221> polyA_site
<222> 673..684
<400> 291
atccccawns cactetetea cagagactgt tetttteett etgagaceet actccagett
gtagttctaa atctgtgatt atgcactgtc tgtcttcctc ttgaggtcag gggccatttc
ttttgttctc tgctatgctc aggacccaga tcaaaggagc tcagtaacta tttacaggcg
tacatcatat gtggaggaca cttatgctgt g atg gcc cca cac aca gct tcc
                                                                     232
                                   Met Ala Pro His Thr Ala Ser
                                       -35
ttt ggg gtc tgt ccc ctg ctc tcc gtt acc cgc gtg gta gcc act gag
                                                                     280
Phe Gly Val Cys Pro Leu Leu Ser Val Thr Arg Val Val Ala Thr Glu
                                    -20
cac tgg ctc ttc ctg gct tca ctc tct ggc atc aaa act tat cag tcc
                                                                     328
His Trp Leu Phe Leu Ala Ser Leu Ser Gly Ile Lys Thr Tyr Gln Ser
           -10
                            - 5
                                             1
tac atc tca gtc ttt tgc aag gtg aca ctt atc tgattaccta attcacacra
                                                                     381
Tyr Ile Ser Val Phe Cys Lys Val Thr Leu Ile
                        10
aggtgttaat ggtggtaatg gcataktatt tattacccca ggggacccak aacggtggta
                                                                     441
tcaaaacata tcattcccca gtggtttaaa actctggtag ctttccargg aatccaaagt
                                                                     501
ggaatccagt ctccttagct gawttcacag ggccccgtct gcacaacttg gcttctgtcg
gettecetan ecetgaette ecaageetta gteateacce teteteceae ecagggetea
gcacagtacc tggaacagtc aagccctcaa taaatgttta ctgagtgcat yaaaaaaaaa
                                                                    681
aaa
                                                                    684
```

```
<210> 292
```

<211> 628

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 75..482

<221> sig_peptide

<222> 75..128

<221> polyA_signal <222> 595..600 <221> polyA site <222> 618..627 <400> 292 60 aagtgagacc gegeggeaac agettgegge tgeggggage tecegtggge geteegetgg ctgtgcaggc ggcc atg gat tcc ttg cgg aaa atg ctg atc tca gtc gca 110 Met Asp Ser Leu Arg Lys Met Leu Ile Ser Val Ala -10 -15 158 atg ctg ggc gca rgg gct ggc gtg ggc tac gcg ctc ctc gtt atc gtg Met Leu Gly Ala Xaa Ala Gly Val Gly Tyr Ala Leu Leu Val Ile Val acc ccg gga gag cgg cgg aag cag gaa atg cta aag gag atg cca ctg 206 Thr Pro Gly Glu Arg Arg Lys Gln Glu Met Leu Lys Glu Met Pro Leu 20 15 254 Gln Asp Pro Arg Ser Arg Glu Glu Ala Ala Arg Thr Gln Gln Leu Leu. 35 ctg gcc act ctg cag gag gca gcg acc acg cag gag aac gtg gcc tgg 302 Leu Ala Thr Leu Gln Glu Ala Ala Thr Thr Gln Glu Asn Val Ala Trp agg aag aac tgg atg gtt ggc ggc gaa ggc gcc acg gga kgt cac 350 Arg Lys Asn Trp Met Val Gly Gly Gly Gly Ala Thr Gly Xaa His cgt gag acc gga ctt gcc tcc gtg ggc gcc gga cct tgg ctt ggg cgc 398 Arg Glu Thr Gly Leu Ala Ser Val Gly Ala Gly Pro Trp Leu Gly Arg 80 85 agg aat cog agg cag ctt tot cot tog tgg goo can ogg aaa ato ogg 446 Arg Asn Pro Arg Gln Leu Ser Pro Ser Trp Ala Xaa Arg Lys Ile Arg 100 95 amc gaa aat wcc atg cca gga ctc tcc ggg gtc ctg tgaactgccg 492 Xaa Glu Asn Xaa Met Pro Gly Leu Ser Gly Val Leu 115 tegggtgage acgtgteece caaaccetgg actgaetget ttaaggteeg caaggeggge cagggccgag acgcgagtcg gatgtggtga actgaaagaa ccaataaaat catgttcctc 628 cammcaaaaa aaaaah

<210> 293

<211> 813

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 50..631

<221> sig_peptide

<222> 50..244

<223> Von Heijne matrix score 8 seq LTLIGCLVTGVES/KI

<221> polyA_signal

<222> 777..782

<221> polyA_site

<222> 801..812

<400> 293 aaggaaagga ttac	tcgagc cttgtt	agaa tcagacatgg	cttcagggg atg c Met G -65	ag gac 58 ln Asp
gct ccc ctg agc Ala Pro Leu Ser -60	Cys Leu Ser	ccg act aag tgg Pro Thr Lys Trp -55	agc agt gtt tct Ser Ser Val Ser -50	tcc 106 Ser
gca gac tca act	gag aag tca	gcc tct gcg gca	ggc acc agg aat Gly Thr Arg Asn -35	ctg 154 Leu
cct ttt cag ttc	tgt ctc cgg	cag gct ttg agg Gln Ala Leu Arg -20	atg aag gct gcg Met Lys Ala Ala	ggc 202 Gly -15
att ctq acc ctc	att ggc tgc	ctg gtc aca ggc	gtc gag tcc aaa Val Glu Ser Lys 1	atc 250 Ile
tac act cgt tgc Tyr Thr Arg Cys	aaa ctg gca	aaa ata ttc tcg	agg gct ggc ctg Arg Ala Gly Leu 15	gac 298 Asp
aat cyg agg ggc Asn Xaa Arg Gly 20	ttc agc ctt Phe Ser Leu 25	gga aac tgg atc	tgc atg gcg tat Cys Met Ala Tyr	tat 346 Tyr
gag age gge tac	aac acc aca	gcc car acg gtc Ala Gln Thr Val	ctg gat gac ggc Leu Asp Asp Gly	agc 394 Ser 50
atc gac tay ggo	atc ttc caa	atc aac agc tto	gcg tgg tgc aga Ala Trp Cys Arg 65	cgc 442 Arg
gga aag ctg aag Gly Lys Leu Lys 70	gag aac aac	cac tgc cay gto	gcc tgc tca gcc Ala Cys Ser Ala 80	ttg 490 Leu
rtc act gat gad	ctc aca gat Leu Thr Asp	gca att atc tgt	gcc arg aaa att Ala Xaa Lys Ile 95	gtt 538 Val
aaa gag aca caa	gga atg aac Gly Met Asn 105	tat tgg caa ggc	tgg aag aaa cay Trp Lys Lys His	tgt 586 Cys
gag ggg aga gad	ctg tcc gas	tgg aaa aaa ggo Trp Lys Lys Gly 125	tgt gag gtt tcc Cys Glu Val Ser	631
taaactggaa ctgg	acccag gatgct	ttgc ascaacgccc	tagggtttgc agtg ccttctcaaa cttg ttaaatgtca amaa	gagagg 751

<211> 778 <212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 154..576

<221> sig_peptide <222> 154..360

<223> Von Heijne matrix score 4.80000019073486 seq MMVLSLGIILASA/SF

```
<221> polyA_signal
<222> 737..742
<221> polyA_site
<222> 763..775
<400> 294
agtaaaaaaa cactggaata aggaagggct gatgactttc agaagatgaa ggtaagtaga
                                                                      60
aaccgttgat gggactgaga aaccagagtk aaaacctctt tggagcttct gaggactcag
                                                                      120
ctggaaccaa cgggcacagt tggcaacacc atc atg aca tca caa cct gtt ccc
                                                                      174
                                     Met Thr Ser Gln Pro Val Pro
aat gag acc atc ata gtg ctc cca tca aat gtc atc aac ttc tcc caa
                                                                      222
Asn Glu Thr Ile Ile Val Leu Pro Ser Asn Val Ile Asn Phe Ser Gln
                           ~~`55
gca gag aaa ccc gaa ccc acc aac cag ggg cag gat agc ctg aag aaa
                                                                      270
Ala Glu Lys Pro Glu Pro Thr Asn Gln Gly Gln Asp Ser Leu Lys Lys
   -45
                       -40
                                           -35
cat cta cac gca gaa atc aaa gtt att ggg act atc cag atc ttg tgt
                                                                     318
His Leu His Ala Glu Ile Lys Val Ile Gly Thr Ile Gln Ile Leu Cys
                   -25
                                        -20
                                                            -15
ggc atg atg gta ttg agc ttg ggg atc att ttg gca tct gct tcc ttc
                                                                     366
Gly Met Met Val Leu Ser Leu Gly Ile Ile Leu Ala Ser Ala Ser Phe
               -10
                                    - 5
tot cca aat tit acc caa gtg act tot aca ctg ttg aac tot gct tac
                                                                     414
Ser Pro Asn Phe Thr Gln Val Thr Ser Thr Leu Leu Asn Ser Ala Tyr
                           10
cca ttc ata gga ccc ttt ttt gtr akt aaa btt tct gag gag ggc agg
                                                                      462
Pro Phe Ile Gly Pro Phe Phe Val Xaa Lys Xaa Ser Glu Glu Gly Arg
                       25
atg ggg caa ara ggg gag gaa rat voc aat agc tta aac tto cca sct
                                                                     510
Met Gly Gln Xaa Gly Glu Glu Xaa Xaa Asn Ser Leu Asn Phe Pro Xaa
                   40
                                       45
gcc agc ttg cta tkt ttg atc tgc cag gav caa gga ttc aac ggt gaa
                                                                     558
Ala Ser Leu Leu Xaa Leu Ile Cys Gln Xaa Gln Gly Phe Asn Gly Glu
               55
                                    60
tot tgt tot cot gtc ggg targataaca ggggttgctt rattttagat
                                                                     606
Ser Cys Ser Pro Val Gly
caatttctta tcagactcaa ataaacattt cttttgaaaa tcatcttatt cttcacatta
                                                                     666
tcatcttgag ctatgatgga aactagtgas ktctctccag gtttaggcga aaaaaaaatc
                                                                     726
                                                                     778
catgaattag gataaagttg ggaaggaaca ttttatacaa aaaaaaaaah cc
```

<210> 295 <211> 1060 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 154..897

<221> sig_peptide <222> 154..360 <223> Von Heijne matrix score 4.80000019073486 seq MMVLSLGIILASA/SF

<221> polyA_signal <222> 1017..1022

*

<221> polyA_site <222> 1044..1054

agta	gttg	aaa d gat g	ggad	tgag	ga aa	accaç	gagtl	c aaa	acct c atq	ctt g aca	tgga a tca	agctt a caa	a cct a cct a Pro	gagga t gtt o Val	agtaga actcag c ccc l Pro	60 120 174
aat Asn	gag Glu	acc Thr	atc Ile	ata Ile	gtg Val	ctc Leu	cca Pro -55	tca Ser	aat Asn	gtc Val	atc Ile	aac Asn -50	-65 ttc Phe	tcc Ser	caa Gln	222
														aag Lys		270
cat His -30	cta Leu	cac His	gca Ala	gar Glu	rtc Xaa -25	aaa Lys	gtt Val	att Ile	Gly aaa	act Thr -20	atc Ile	cag Gln	atc Ile	ttg Leu	tgt Cys -15	318
ggc	atg Met	atg Met	gta Val	ttg Leu -10	agc Ser	ttg Leu	gly aaa	atc Ile	att Ile -5	ttg Leu	gca Ala	tct Ser	gct Ala	tcc Ser 1	ttc Phe	366
														gct Ala		414
														tca Ser		462
gcc Ala 35	aca Thr	aaa Lys	aaa Lys	agg Arg	tta Leu 40	acc Thr	aac Asn	ctt Leu	ttg Leu	gtg Val 45	cat His	acc Thr	acc Thr	ctg Leu	gtt Val 50	510
gga Gly	agc Ser	att Ile	ctg Leu	agt Ser 55	gct Ala	ctg Leu	tct Ser	gcc Ala	ctg Leu 60	gtg Val	ggt Gly	ttc Phe	att Ile	ayc Xaa 65	ctg Leu	558
														gag Glu		606
Xaa	Lys	Asn 85	Asn	Ile	Pro	Thr	Xaa 90	Xaa	Tyr	Val	Xaa	Tyr 95	Phe	tat Tyr	His	654
Asp	Ser 100	Leu	Tyr	Thr	Thr	Asp 105	Xaa	Tyr	Thr	Ala	Lys 110	Ala	Xaa	ctg Leu	Ala	702
Gly 115	Thr	Leu	Ser	Leu	Met 120	Leu	Ile	Cys	Thr	Leu 125	Leu	Glu	Phe	tgc Cys	Xaa 130	750
Xaa	Val	Leu	Thr	Ala 135	Val	Leu	Arg	Trp	Lys 140	Gln	Ala	Tyr	Ser	gac Asp 145	Phe	798
														tct Ser		846
														ttg Leu		894
tct Ser	taa	gaaa	aaa q	ggga	gaaat	a tt	caato	cagaa	a agt	tgai	tct	tate	gataa	ata		947
			aacca ttaaa												ttaaa	1007 1060

```
<210> 296
<211> 444
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 146..292
<221> sig_peptide
<222> 146..253
<223> Von Heijne matrix
     score 5.5
     seq FTSMCILFHCLLS/FQ
<221> polyA_signal
<222> 395..400
<221> polyA site
<222> 433..444
<400> 296
aacttgggac aagaratcaa actttaaaga tggtctaaag cccctcttaa aggtctgact
gtgtcggacc tctagagcta atctcactag atgtgagcca ttgtttatat tctagccatc
ctttcatttc attctagaag acccc atg caa gtt ccc cac cta agg gtc tgg
                         Met Gln Val Pro His Leu Arg Val Trp
                              -35
aca cag gtg awa gat acc ttc att ggt tat aga aat ttg gga ttt aca
Thr Gln Val Xaa Asp Thr Phe Ile Gly Tyr Arg Asn Leu Gly Phe Thr
       -25
                          -20
agt atg tgc ata ttg ttc cac tgt ctt ctt agc ttt cag gtt ttc aaa
Ser Met Cys Ile Leu Phe His Cys Leu Leu Ser Phe Gln Val Phe Lys
                                               5
aag aaa aga aaa ctt ara ctt ttc tgatgttctt ttttacgtaa ataaccattt
Lys Lys Arg Lys Leu Xaa Leu Phe
               10
tattgttgtt ttgctttttc tgccttcaaa ctactcccac aggccaaata tavctggctg
aa
<210> 297
<211> 754
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 126..383
<221> sig_peptide
<222> 126..167
<223> Von Heijne matrix
     score 7.5
     seq VALNLILVPCCAA/WC
<221> polyA signal
<222> 726..731
```

<221> polyA_site <222> 743..754

60

120 172

220

268

322

382

442

444

<400> 297	
aattgtatgt tacgatgttg tattgatttt taagaaagta attkratttg taaaacttct gctcgtttac actgcacatt gaatacaggt aactaattgg wwggagaggg gaggtcactc ttttg atg gtg gcc ctg aac ctc att ctg gtt ccc tgc tgc gct gct tgg Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp	60 120 170
tgt gac cca cgg agg atc cac tcc cag gat gac gtg ctc cgt agc tct  Cys Asp Pro Arg Arg Ile His Ser Gln Asp Asp Val Leu Arg Ser Ser  10 15	218
gct gct gat act ggg tct gcg atg cag cgg cgt gag gcc tgg gct ggt Ala Ala Asp Thr Gly Ser Ala Met Gln Arg Arg Glu Ala Trp Ala Gly 20 25 30	266
tgg aga agg tca caa ccc ttc tct gtt ggt ctg cct tct gct gaa aga Trp Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg 35 40 45	314
ctc gag aac caa cca ggg aag ctg tcc tgg agg tcc ctg gtc gga gag Leu Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu 50 55 60 65	362
gga cat aga atc tgt gac ctc tgacrrctgt gaasccaccc tgggctacar Gly His Arg Ile Cys Asp Leu 70	413
aaaccacagt cttcccagca attattacaa ttcttgaatt ccttggggat tttttactgc cctttcaaag cacttaaktg tkrratctaa cgtkttccag tgtctgtctg aggtgactta aaaaatcaga acaaaacttc tattatccag agtcatggga gagtacaccc tttccaggaa taatgttttg ggaaacactg aaatgaaatc ttcccagtat tataaattgt gtatttaaaa aaaagaaact tttctgaatg cctacctggc ggtgtatacc aggcagtgtg ccagtttaaa aagatgaaaa agaataaaaa cttttgagga aaaaaaaaaa	473 533 593 653 713 754
<211> 629 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 66497	
<221> sig_peptide <222> 66239 <223> Von Heijne matrix score 5.40000009536743 seq QLLDSVLWLGALG/LT	
<221> polyA_signal	
<221> polyA_site <222> 618629	
<pre>&lt;400&gt; 298 aactcccaga atgctgacca aagtgggagg agcactaggt cttcccgtca cctccacctc tctcc atg acc cgg ctc tgc tta ccc aga ccc gaa gca cgt gag gat ccg Met Thr Arg Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro</pre>	60 110
atc cca gtt cct cca agg ggc ctg ggt gct ggg gag ggg tca ggt agt  Ile Pro Val Pro Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser  -40 -35 -30	158
cca gtg cgt cca cct gta tcc acc tgg ggc cct agc tgg gcc cag ctc Pro Val Arg Pro Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu	206

-25 -20 -15	
ctg gac agt gtc cta tgg ctg ggg gca cta gga ctg aca atc cag gc Leu Asp Ser Val Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln A	la
gtc ttt tcc acc act ggc cca gcc ctg ctg ctg ctt ctg gtc agc tt Val Phe Ser Thr Thr Gly Pro Ala Leu Leu Leu Leu Leu Val Ser Ph	tc 302
ctc acc ttt gac ctg ctc cat agg ccc gca gtc aca ctc tgc cac ag Leu Thr Phe Asp Leu Leu His Arg Pro Ala Val Thr Leu Cys His So	gc 350 er
gca aac ttc tca cca ggg gcc aga gtc agg ggg ccg gtg aag gtc cc Ala Asn Phe Ser Pro Gly Ala Arg Val Arg Gly Pro Val Lys Val Lo	tg 398 eu
gac age agg agg etc tac tec tge aaa tgg gta eag tet eag gac ag Asp Ser Arg Arg Leu Tyr Ser Cys Lys Trp Val Gln Ser Gln Asp A	ac 446 sn
tta gcc tcc agg aag cac tgc tgc tgc tca tgg ggc tgg gcc c Leu Ala Ser Arg Lys His Cys Cys Cys Cys Ser Trp Gly Trp Ala A	rg
70 75 80 tcc tgaaaacctg tggcatgccc ttgwaccctg cttggcctgg ctttctgcct	547
Ser	gacc 607
ccatccttgg gcctgakanc ccctccccac aactcagtgt ccttcaaata tacaat	629
<210> 299	
<211> 765	
<212> DNA <213> Homo sapiens	
<220>	
<221> CDS <222> 49411	
7000 3711311	
<pre>&lt;221&gt; sig_peptide &lt;222&gt; 4996</pre>	
<223> Von Heijne matrix	
score 10.1000003814697 seq LVLTLCTLPLAVA/SA	
sed nontreprovious	
<221> polyA_signal <222> 732737	
<221> polyA_site <222> 750763	
<400> 299	
aaagatccct gcagcccggc aggagagaag gctgagcctt ctggcgtc atg gag Met Glu -15	agg 57 Arg
ctc gtc cta acc ctg tgc acc ctc ccg ctg gct gtg gcg tct gct g	ggc ⁻ 105
Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala Ser Ala G	31A
tgc gcc acg acg cca gct cgc aac ctg agc tgc tac cag tgc ttc a Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln Cys Phe I 5 10 15	aag 153 Lys
qtc agc agc tgg acg gag tgc ccg ccc acc tgg tgc agc ccg ctg g	gac 201
Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser Pro Leu A	Asp :
caa gto too ato too aac gag gtg gtc gtc tot tit agt gag toy o	cc 249
Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Ser Glu Ser F	Pro

• •	
40 45 50	
ccg ggc aga ggg cas gtg cca bgt gcc ggg gaa kgg ccg gtg ccc ccg Pro Gly Arg Gly Xaa Val Pro Xaa Ala Gly Glu Xaa Pro Val Pro Pro 55 60 65	297 .
cct ctc wkc gac tta bct atg act cct cgg ckc ycc agg gcc tgg ggc Pro Leu Xaa Asp Leu Xaa Met Thr Pro Arg Xaa Xaa Arg Ala Trp Gly	345
70 75 80  cck gtg ggt ccd aaa gtg cct cct gct gtc tct ccc gcg ctg ggc tcg  Pro Val Gly Pro Lys Val Pro Pro Ala Val Ser Pro Ala Leu Gly Ser	393
85 90 95 ggc gag cat ccs rva btg tgaatkkkga cttttttctc ckccatttga Gly Glu His Pro Xaa Xaa	441
100 105 agtgtcacta ggaactgtca gcaggacaaa ggctctgatg tcactgaatt tacaaaraca	501
gcaggaacrs ackggtgggg atgggcagct gttcrarger atgggtkate tgccetteet	561
ggcacagcac artacacctg ccatacaacc carcatcagg cakgctgcac tggaatcgat acagtgtatg acaatgtcat atagtataac acaacataat gaatataacg tgtatattgc	621 681
accttaatat aatacgatgt aatataatgc tacataatac aacataatat aataaaatag	741
aatgcaacac aaaaaaaaa aacc	765
220. 200	
<210> 300 <211> 623	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<221> sig_peptide <222> 4996	
<223> Von Heijne matrix	
score 10.100003814697 seq LVLTLCTLPLAVA/SA	
<221> polyA_signal	
<222> 593598	
<221> polyA_site	
<222> 612623	
<400> 300	
aaagatccct gcagcccggc aggagagaag gctgagcctt ctggcgtc atg gag agg Met Glu Arg -15	57
ctc gtc cta acc ctg tgc acc ctc ccg ctg gct gtg gcg tct gct ggc	105
Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala Ser Ala Gly	
-10 -5 1 tgc gcc acg acg cca gct cgc aac ctg agc tgc tac cag tgc ttc aag	153
Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln Cys Phe Lys	
5 10 15 gtc agc agc tgg acg gag tgc ccg ccc acc tgg tgc agc ccg ctg gac	201
Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser Pro Leu Asp	
20 25 30 35	249
caa gtc tgc atc tcc aac gag gtg gtc gtc tct ttt aaa tgg agt gta Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Lys Trp Ser Val	
40 45 50	207
cgc gtc ctg ctc agc aaa cgc tgt gct ccc aga tgt ccc aac gac aac Arg Val Leu Leu Ser Lys Arg Cys Ala Pro Arg Cys Pro Asn Asp Asn	297
55 60 65	

atg aa	ak t	tc g	aa t	gg 1	tcg	ccg	gcc	ccc	atg	gtg	caa	ggc	gtg	atc	acc	345
Met Xa	aa P	he G	lu I	rp :	Ser	Pro	Ala	Pro	Met	Val	Gln	GTA	Val	Ile	Thr	•
	7	0					75					80				3.93
agg cg	gc t	gc t	gt t	cc	tgg	gct	ctc	tgc	aac	agg	gca	ctg	acc	D~	cag	3.33
Arg Ar	rg C	ys C	ys S	Ser '	Trp	Ala	Leu	Cys	Asn	Arg	Ala	Leu	THE	PIO	GIII	
85	5					90					95		~	cct	tca	441
gag gg	gg c	gc t	gg g	acc	ctg	cra	agg	999	CLC	tau	Tou	Gln	Asn	Pro	Ser	
Glu Gl	ly A	rg 1	rp A			Xaa	GIY	GIY	Leu	110	Den	GIII	voh	110	115	
100					105					110		ctc	cca	ctc		489
agg gg	gc a	ra a	aaa a	acc	tgg	gtg	cgg	Dro	cay	Len	999 999	Len	Pro	Leu	Cvs	
Arg G	ly X	aa I			Trp	vaı	Arg	PLO	125	пеп	Gry			130		
ctt co				120		ctc	tac	cca		gaa	acc	cag	qaa	gga		534
Leu Pi	cc a	we t	200 8	aac Nan	200	Len	Cve	Pro	Xaa	Glu	Thr	Gln	Glu	Gly		
Leu Pi	ro v		135	A511	FIU	пси	Cys	140	••••				145	•		
+	a+ a+	~ ~	150	~~~	a cc	rate	cati		acca	acra	ctt	cacc	ctc	tgg	aracaa	594
taacac	tata	9 95	geger	ccaa	a aa	12222	aaaa	- 55.	,							623
tadacı		a ç	3000	ccaa												
<210>	301															
<211>																
<212>																
<213>			apie	ns												
			•													
<220>																
<221>	CDS	3														
<222>	86.	41	5													
<221>				.e												
<222>																
	. Voi	n He	iine	mat	rix											
<223>			-)				_									
<223>	SC	ore	9.80	0000	190	7348	6									
<223>	SC	ore	9.80 IGLT	0000	190	7348	6									
	sed	ore q FT	9.80 IGLT	LLL(	190	7348	6									
<221>	sco seo	ore q FT lyA_	9.80 IGLT sign	LLL(	190	7348	6									
	sco seo	ore q FT lyA_	9.80 IGLT sign	LLL(	190	7348	6									
<221> <222>	sec sec po:	ore q FT lyA_ 05	9.80 IGLT sign 45	OOOO	190	7348	6									
<221> <222>	sco seo po: 54	ore q FT lyA_ 05	9.80 IGLT sign 45 site	OOOO	190	7348	6									
<221> <222>	sco seo po: 54	ore q FT lyA_ 05	9.80 IGLT sign 45 site	OOOO	190	7348	6									
<221><222><222><221><222>	scc sec po: 54 po: 56 s 30	ore TFT lyA_ 05 lyA_ 05	9.80 IGLT sign 45 site 71	OOO( LLL( aal	)190 GXQA	7348 /MP										60
<221><222><222><221><222><400>	scc sec sec sec sec sec sec sec sec sec	ore TFT lyA_ 05 lyA_ 05	9.80 IGLT sign 45 site 71	OOO(	0190 GXQA ta t	7348 /MP	attt	a aç	gaagc	atco	t tot	:gcca	aaga	ccaa	aaaggaa	60
<221><222><222><221><222><400>	scc sec sec sec sec sec sec sec sec sec	ore TFT lyA_ 05 lyA_ 05	9.80 IGLT sign 45 site 71	OOO(	0190 GXQA ta t	7348 /MP	attt	ara	ctg	ato	g gta	a ctt	gu		aaaggaa cacc	60 <b>112</b>
<221><222><222><221><222>	scc sec sec sec sec sec sec sec sec sec	ore TFT lyA_ 05 lyA_ 05	9.80 IGLT sign 45 site 71	OOO(	0190 GXQA ta t	7348 /MP	attt	ara	ctg	ato	g gta	l Lei	ı Val		aaaggaa : acc : Thr	
<221><222><221><222><400>aaaaaaaaaaaaaaaaaaaaaaa	scc sec po: 54 po 56 30 actc	ore q FT lyA_ 05 lyA_ 1 ac c aa b	9.80 IGLT sign 45 site 71	oooc aal	D190 JXQA tg t ag c	7348 /MP gago caaa	attt atc Met	ara Xaa	ctg Leu	Met	g gta : Val	l Lev	yul Val	. Phe	Thr	112
<221><222><222><221><222><400>aaaaa	scc sec po: 54 po 56 30 actc	ore q FT lyA_ 05 lyA_ 05 lac caa b	9.80 IGLT sign 45 site 71 cagt	COOC LLLC	tg tag c	7348 /MP gago caaa	attt atg Met -20	ara Xaa Xaa	ctg Leu caa	Met	g gta : Val	l Lev -15	yal Val	Phe aat	Thr	
<221><222><222><221><222><400>aaaaa	scc sec po: 54 po 56 30 actc	ore q FT lyA_ 05 lyA_ 05 lac caa b	9.80 IGLT sign 45 site 71 cagt	COOC LLLC	tg tag c	7348 /MP gago caaa	attt atg Met -20	ara Xaa Xaa	ctg Leu caa	Met	g gta : Val : atg a Met	l Lev -15	yal Val	Phe aat	Thr cgc	112
<221><222><222><221><222><400>aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	score second sec	ore q FT lyA_ 05 lyA_ ac c aa b cta Leu	9.80 IGLT sign 45 site 71 cagto	ttg	tg tag c	7348 /MP gagc caaa Leu -5	attt atg Met -20 a gga a Gly	y ara : Xaa ) a rtt / Xaa	ctg Leu caa Gln	Met Met GCG	y gta : Val : atg a Met	Lev -15 g cct	Val	Phe a aat a Asr	Thr cgc Arg 5	112
<221><222><222><221><222><400>aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	score secore sec	ore I FT  IYA_ 05  IYA_ 1	9.80 IGLT sign 45 site 71 cagto	ttg	tg tag c	7348 /MP gagc caaa Leu -5	attt atg Met -20 1 ggs 1 Gly	ara Xaa Xaa Xaa Xaa	ctg Leu caa Gln gat	Met Met GCG Ala	y grade Value atom American Am	l Let -15 g cct t Pro	Val	Phe a aat Asr	Thr cgc Arg 5	112
<221><222><222><221><222><400>aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	score secore sec	ore I FT  IYA_ 05  IYA_ 1	9.80 IGLT sign 45 site 71 cagto	ttg	tg tag c	7348 /MP gagc caaa Leu -5	attt atg Met -20 1 ggs 1 Gly	ara Xaa Xaa Xaa Xaa	ctg Leu caa Gln gat Asp	Met Met GCG Ala	y grade Value atom American Am	l Let -15 g cct t Pro	Val	Phe a aat Asr a Asr	Thr cgc Arg 5	112
<221><222><222><221><222><400>aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	scc sec sec sec sec sec sec sec sec sec	ore I FT  IYA_ 05  IYA_ ac c aa b  cta Leu tgc Cys	9.80 IGLT sign 45 site 71 cagto	ttg Leu aga Arg	tg tag c	gago caaa Leu -5 ata	attt atc Met -20 gga Gly	xaa Xaa Xaa Xaa Xaa Aaaa Lys	caa Caa Gln gat Asp	Met gco Ala	val c ato a Met l c aao	Level	Val Segration Ala	Phe aat Asr aac Asr 20	Thr cgc Arg 5 ctt Leu	112 160 208
<221><222><221><222><400>aaaaa agaagatte t	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_ 0 5  I YA_ 1 ac c aa b  cta Leu tgc Cys	9.80 IGLT sign 45 site 71 cagto act Thr tac	ttg aga Leu aga Arg	tg tag c	7348 /MP gaggaaa cta Leu -5 ata Ile	attt atc Met -20 gga Gly	ara Xaa Xaa Xaa Xaa Aaaa Lys	ctg Leu caa Gln gat S Asp 15	Met gco Ala cac His	y grade Value atom Assist grade	Levi -15 g cott Pro	yali Vali Vali Vali Vali Vali Vali Vali V	Phe aat Asr aac 20 cag	Thr cgc Arg cctt Leu ggat	112
<221><222><221><222><400>aaaaa agaagatte t	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_ 0 5  I YA_ 1 ac c aa b  cta Leu tgc Cys	9.80 IGLT sign 45 site 71 cagt ggcc act Thr tac Tyr gta Val	ttg aga Leu aga Arg	tg tag c	7348 /MP gaggaaa cta Leu -5 ata Ile	attt atc Met -20 gga Gly	xaa Xaa Xaa Xaa Xaa Aaaa Lys Aaaa Cag	ctg Leu caa Gln gat S Asp 15	Met gco Ala cac His	y grade Value atom Assist grade	Levi -15 g cott Pro	yali Valis gca o Ala cac His	Phe aat Asr aac 20 cag	Thr cgc Arg 5 ctt Leu	112 160 208
<221><222><221><222><400>aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	scc sec sec sec sec sec sec sec sec sec	ore I FT  IYA_ 05  IYA_ 05  Lac b  cta Leu  tgc Cys  gga Gly	9.80 IGLT sign 45 site 71 cagco act Thr tac Tyr gta 25	gaggaaaattg Leu aga Arg	tg tcag ctg Leu aacs	gaggaaaa Leu -5 ata Ile	attt atg Met -20 gga Gly cta Lev	xaa Xaa Xaa Xaa Xaa Aaa Lys Aaaa Cag T Gli	ctg Leu caa Gln agat Asp 15 atti	Met gco Ala cac His gat	g gta Val c ato a Met l c aac s Asi t gto	t Cttl Let -15 g cct t Pro c tgt n Cys c aat	yali Valis gca para para para para para para para pa	Phe aat Asr aac aac 20 cac Cac	Thr cgc Arg cctt Leu ggat Asp	112 160 208 256
<221> <222> <221> <222> <400> aaaaa agaag att g Ile G ctc t Leu S crog g	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_ 0 5  I YA_ 1 ac b  cta Leu tgc Cys gga Gly	9.80 IGLT sign 45 site 71 caggco act Thr tac Tyr gtal 25 at	gaggaaaattg Leu aga Arg gct Ala	tag ctg Leu aags Lys Asp	gaga caaa cta Leu -5 ata Ile	attt atg	ara Xaa Xaa Xaa Xaa Aaaa Lys Gli 30	ctg Leu caa Gln agat Asp 15 g att	Met gcc Alac cac His gate Asp	g gta Val c ato a Met l c aac Ass t gto p Val	t tact	yali Vali Vali Vali Vali Vali Vali Vali V	Phe aat Asr 20 cag Glr	Thr  cogc Arg 5 cott Leu ggat Asp	112 160 208
<221> <222> <221> <222> <400> aaaaa agaag att g Ile G ctc t Leu S crog g	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_  0 5  I YA_  1 C C C C C C C C C C C C C C C C C C	9.80 IGLT sign 45 site 71 caggco act Thr tac Tyr gtal 25 at	gaggaaaattg Leu aga Arg gct Ala	tag ctg Leu aags Lys Asp	gaga caaa cta Leu -5 ata Ile	Attt Att Met -20 Gga Gly Cta Lev Act Thi	ara Xaa Xaa Xaa Xaa Aaaa Lys Gli 30	ctg Leu caa Gln agat Asp 15 g att	Met gcc Alac cac His gate Asp	g gta Val c ato a Met l c aac Ass t gto p Val	t tack	yali Vali Vali Vali Vali Vali Vali Vali V	Phe aat Asr 20 cag Glr	Thr cgc Arg cctt Leu ggat Asp	112 160 208 256
<221> <222> <221> <222> <400> aaaaa agaag  att g  tle g  ctc t  Leu s  craft  His I	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_ 0 5  I YA_ 1 C C A A A A A A A A A A A A A A A A A	9.80 IGLT sign 45 site 71 caggco act Thr tac Tyr gtal 25 gat Asp	ttg gag: aaaa: Leu aga Arg 10 gct Ala	tg tcag ctag Lys	gago caaa Leu -5 ata Ile Cto Lev	attt atc Met -20 gga Gly cta Lev g aca Thi	xaa Xaa Xaa Xaa Xaa Aaaa Lys Aaaaa Cag T Gli 30 t gag	ctg Leu caa Gln agat 15 gatt 11e	Met gcc Ala Cac His gate Asp	c atomic and the control of the cont	t tacks Ty:	t cac t gto t gto t gto t gto t gto t gto c tgo c tgo	Phe aat Asr 20 cag Glr Glr aac Asr 3 Asr 3 Cag	Thr  c cgc Arg 5 c ctt Leu g gat n Asp c ttc n Phe	112 160 208 256
<221> <222> <221> <222> <400> aaaaa agaag  att g  tle G  ctc t  Leu S  crog g  Pro G  cat t  His I	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_  0 5  I YA_  1 ac b  cta  tgc  Cys  ggly  tgg  Trp  40	9.80 IGLT sign 45 site 71 caggco act Thr tac Tyr gtal 25 t Asp	ttg gaga aaaa Leu aga Arg gct Ala	tag to Leu aags Lys	gagaaaa cta Leu -5 ata Ileu E Gly	attt atc Met -20 gga Gly cta Let Thi ty 45	g arage arage Xaaa Xaaa Xaaa Aaaa Aaaa Aaaa Aaaa Aaa	ctg Leu caa Gln gat 15 att gatg att gatg	Met gcc Ala cac His gate Asp	e atomic and the control of the cont	t tacks to test	yali Vali E gca D Ala C cac S His T yal C tgc C tgc C tgc	Phe aat Asr 20 Cag Cag Asr Asr Asr Caac	Thr  cogc Arg cott Leu gat Asp cottc Asp	112 160 208 256 304
<221><222><222><222><400>aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_  0 5  I YA_  1 ac b  cta  tgc  Cys  ggly  tgg  Trp  40	9.80 IGLT sign 45 site 71 caggco act Thr tac Tyr gtal 25 t Asp	ttg gaga aaaa Leu aga Arg gct Ala	tag to Leu aags Lys	gago caaa Leu -5 ata Ile S Ile S Gly	attt atc Met -20 gga Gly cta Let Thi ty 45	g arage arage Xaaa Xaaa Xaaa Aaaa Aaaa Aaaa Aaaa Aaa	ctg Leu caa Gln gat 15 att gatg att gatg	Met gcc Ala cac His gate Asp	g gta Val c ato a Met c aao s Asi t gto p Val c tg e Cy	t tack to the text	yali Vali E gca D Ala C cac S His T yal C tgc C tgc C tgc	Phe aat Asr 20 Cag Cag Asr Asr Asr Caac	Thr  c cgc Arg 5 c ctt Leu g gat n Asp c ttc n Phe	112 160 208 256 304
<pre>&lt;221&gt; &lt;222&gt; &lt;221&gt; &lt;222&gt; &lt;400&gt; aaaaa agaag  att g  Ile G  ctc t  Leu S  craft  August  August  Lys G  August  Lys G  August  Augus</pre>	scc sec sec sec sec sec sec sec sec sec	ore I FT  I YA_ 0 5  I yA_ 1 ac b  cta Leu tgc Cys ggly tgg Trp 40 atte	9.80 IGLT sign 45 site 71 caggco act Thr tac Tyr gtal 25 at Asp gct	ttg gaga caaa ttg Leu aga Arg gct Ala ggg Ctg	tag ctg Leu aag Lys gac Lys tag Lys	gagaaaa cta Leu -5 ata Ile Ctc Leu Gall Ccc	attt atc Met -20 gga Gly cta Let Thi ty 45 caas	xaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	ctg Leu caa Gln agat 15 att att u Met	Met Met Good Alac Cac His gate Asp atc Ilc	t gtale atom and a determination of the control of	t tack to the track to the trac	yali Fin Validation Alamanda A	Phe aata Ass 20 cag Cag Cag Ass Ass Ass Ass Ass Ass Ass Ass Ass As	Thr  cogc Arg cott Leu gat Asp cottc Asp	112 160 208 256 304

Leu Phe Arg Asp Ser Leu Gln Gln Ser Met Arg Ile Phe Met Tyr Ser 70 75 80 85 ggc gaa cac cat tcc tgatttccca caaactgcac tacatcagta taactgcatt	455
Gly Glu His His Ser	
tctagtttct atatagtgca atagagcata gattctataa attcttactt gtctaagaaa gtaaatctgt gttaaacaag tagtaataaa agttaattca atccaaaaaa aaaaaa	515 571
<210> 302 <211> 612	
<212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 56268	
<pre>&lt;221&gt; sig_peptide &lt;222&gt; 56100 &lt;223&gt; Von Heijne matrix</pre>	
<221> polyA_signal <222> 584589	
<221> polyA_site <222> 601612	
<400> 302 ctaatcgaaa agggggattt teeggtteeg geetggegag agtttgtgeg gegae atg Met -15	58
aaa ctg ctt acc cac aat ctg ctg agc tcg cat gtg cgg ggg gtg ggg Lys Leu Leu Thr His Asn Leu Leu Ser Ser His Val Arg Gly Val Gly -10 -5	106
tcc cgt ggc ttc ccc ctg cgc ctc cag gcc acc gag gtc cgt atc tgc Ser Arg Gly Phe Pro Leu Arg Leu Gln Ala Thr Glu Val Arg Ile Cys	154
cct gtg gaa ttc aac ccc aac ttc gtg gcg cgt atg ata cct aaa gtg Pro Val Glu Phe Asn Pro Asn Phe Val Ala Arg Met Ile Pro Lys Val	202
gag tgg tcg gcg ttc ctg gag gcg rmc gat aac ttg cgt ctg atc cag Glu Trp Ser Ala Phe Leu Glu Ala Xaa Asp Asn Leu Arg Leu Ile Gln	250
gtg ccg aga agg gcc ggt tgagggatat gaggagaatg aggagtttct Val Pro Arg Arg Ala Gly	298
gaggaccatg caccacctgc tgctggaggt ggamstgaka gagggcaccc tgcagtgccc ggaatctgga cgtatgttcc ccatcagccg cgggatcccc aacatgctgc tgagtgaaga ggaaactgag agttgattgt gccaggcgcc agtttttctt gttatgactg tgtatttttg ttgatctata ccctgtttcc gaattctgcc gtgtgtatcc ccaacccttg acccaatgac accaaacaca gtgtttttga gctcggtatt atatatttt ttctcattaa aggtttaaaa ccaaaaaaaa aaaa	358 418 478 538 598 612

<211> 539 <212> DNA

```
<213> Homo sapiens
<220>
<221> CDS
<222> 32..328
<221> sig peptide
<222> 32..103
<223> Von Heijne matrix
     score 4.59999990463257
     seq FFIFCSLNTLLLG/GV
<221> polyA_signal
<222> 508..513
<221> polyA_site
<222> 528..539
<400> 303
                                                                       52
aacaactate etgeetgetg ettgetgeae e atg aag tet gee aag etg gga
                                  Met Lys Ser Ala Lys Leu Gly
                                                    -20
                                                                      100
ttt ctt cta aga ttc ttc atc ttc tgc tca ttg aat acc ctg tta ttg
Phe Leu Leu Arg Phe Phe Ile Phe Cys Ser Leu Asn Thr Leu Leu Leu
                            -10
                                                                      148
ggt ggt gtt aat aaa att gcg gag aag ata tgt gga gac ctc aaa gat
Gly Gly Val Asn Lys Ile Ala Glu Lys Ile Cys Gly Asp Leu Lys Asp
                                                             15
                    5
ccc tgc aaa ttg gac atg aat ttt gga agc tgc tat gaa gtt cac ttt
                                                                      196
Pro Cys Lys Leu Asp Met Asn Phe Gly Ser Cys Tyr Glu Val His Phe
                20
aga tat ttc tac aac aga acc tcc aaa aga tgt gaa act ttt gtc ttc
Arg Tyr Phe Tyr Asn Arg Thr Ser Lys Arg Cys Glu Thr Phe Val Phe
            35
                                40
tcc agc tgt aat ggc aac ctt aac aac ttc aag ctt aaa ata gaa cgt
                                                                      292
Ser Ser Cys Asn Gly Asn Leu Asn Asn Phe Lys Leu Lys Ile Glu Arg
                                                60
        50
                            55
gaa gta kcc tgt gtt gca aaa tac aaa cca ccg agg tgagaggatg
                                                                      338
Glu Val Xaa Cys Val Ala Lys Tyr Lys Pro Pro Arg
                        70
tgaactcatg aagttgtctg ctgcaccatc cgaaataaag acacaagaaa attcaractg
                                                                      398
atttwgaaat ctttgttwta tttccmymak ggcgwktaag cttccatatg tttgctattt
                                                                      458
tectgacect agttttgtet tteetggaaa ttaactgtat gakeattasa atgaaagagt
                                                                      518
                                                                      539
ctttctgtca aaaaaaaaa a
<210> 304
```

```
<211> 964

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 21..527

<221> sig_peptide

<222> 21..95

<223> Von Heijne matrix

score 8.5

seq LKVLLLPLAPAAA/QD
```

<221> polyA_signal <222> 921..926

<221> polyA_site <222> 953..963

<400> 304 agggcggatc ttctccggcc atg agg aag cca gcc gct ggc ttc ctt ccc tca 53 Met Arg Lys Pro Ala Ala Gly Phe Leu Pro Ser -20 -25 101 ctc ctg aag gtg ctg ctc ctg cct ctg gca cct gcc gca gcc cag gat Leu Leu Lys Val Leu Leu Pro Leu Ala Pro Ala Ala Ala Gln Asp -5 -10 teg act cag gee tee act eca gge age eet ete tet eet ace gaa tae 149 Ser Thr Gln Ala Ser Thr Pro Gly Ser Pro Leu Ser Pro Thr Glu Tyr 10 caa ege tte tte gea etg etg act eea ace tgg aag gea gar act ace 197 Gln Arg Phe Phe Ala Leu Leu Thr Pro Thr Trp Lys Ala Glu Thr Thr 25 tgc cgt ctc cgt gca acc cac ggc tgc cgg aat ccc aca ctc gtc cag 245 Cys Arg Leu Arg Ala Thr His Gly Cys Arg Asn Pro Thr Leu Val Gln 40 ctg gac caa tat gaa aac cac ggc tta gtg ccc gat ggt gct gtc tgc 293 Leu Asp Gln Tyr Glu Asn His Gly Leu Val Pro Asp Gly Ala Val Cys 60 55 341 tee aac etc cet tat gee tee tgg ttt gag tet tte tge cag ttc act Ser Asn Leu Pro Tyr Ala Ser Trp Phe Glu Ser Phe Cys Gln Phe Thr 75 cac tac cgt tgc tcc aac cac gtc tac tat gcc aag aga gtc ctg tgt 389 His Tyr Arg Cys Ser Asn His Val Tyr Tyr Ala Lys Arg Val Leu Cys 90 tee cag cea gte tet att etc tew eet aac act etc aag gag ata gaa 437 Ser Gln Pro Val Ser Ile Leu Ser Pro Asn Thr Leu Lys Glu Ile Glu 105 sct toa got gaa gto toa coo aco aca gat gao etc coo cat etc aco 485 Xaa Ser Ala Glu Val Ser Pro Thr Thr Asp Asp Leu Pro His Leu Thr 120 125 cca ctt cac agt gac aga acg cca gac ctt cca gcc ctg gcc 527 Pro Leu His Ser Asp Arg Thr Pro Asp Leu Pro Ala Leu Ala 140 135 tgagaggctc agcaacaacg tggaagagct cctacaatcc tccttgtccc tgggaggcca 587 ggagcaagcg ccagagcaca agcaggagca aggagtggag cacaggcagg agccgacaca 707 agaacacaag caggaagagg ggcagaaaca ggaagagcaa gaagaggaac aggaagagga gggaaagcag gaagaaggac aggggactaa ggagggacgg gaggctgtgt ctcagctgca 767 gacagactca gagoccaagt ttoactotga atototatot totaaccott cotottttgo tccccgggta cganaagtag agtctactcc tatgataatg gagaacatcc aggagctcat 947 tcgatcagcc caggaaatag atgaaatgaa tgaaatatat gatgagaact cctactggag 964 aaaccaaaaa aaaaaak

<210> 305

<211> 684

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 147..647

<221> sig_peptide <222> 147..374 <223> Von Heijne matrix score 3.5 seq LASASELPLGSRP/AP

<221> polyA_site <222> 668..681

<400> 305 aacttcctgt gagc acgacttctc ctrg agcggcaaga tcgc	rarmcc ccgact	tgagg cggagad c atg ggc gad	gaa ggtgctgcag	gc ggt tac 173 gg Gly Tyr
cgc atg ctg ggc Arg Met Leu Gly -65	Glu Thr Cys	Ala Asp Cys	Gly Thr lie Let	i Leu Gin
gac aaa cag cgg Asp Lys Gln Arg	Lys Ile Tyr -45	Cys Val Ala	Cys Gln Glu Let	asp Ser
gac gtg gat aaa	gat aat ccc Asp Asn Pro	gct ctg aat Ala Leu Asn	gcc cag gct gcc Ala Gln Ala Ala -25	c ctc tcc 317 a Leu Ser -20
caa get egg gag	cac cag ctg	gcc tca gcc Ala Ser Ala -10	tca gag ctc cc Ser Glu Leu Pr	c ctg ggc 365 o Leu Gly -5
tct cga cct gcg Ser Arg Pro Ala	ccc caa ccc	cca gta cct Pro Val Pro 5	cgt ccg gag ca Arg Pro Glu Hi 10	tgt gag 413 s Cys Glu
gga gct gca gca Gly Ala Ala Ala 15	a gga ctc aag a Gly Leu Lys 20	gca gcc cag Ala Ala Gln	ggg.cca cct gc Gly Pro Pro Al 25	t cct gct 461 a Pro Ala
gtg cct cca aat	aca rat gto Thr Xaa Val	atg gcc tgc Met Ala Cys	aca cag aca gc Thr Gln Thr Al 40	a Leu Leu 45
caa aag ctg acg	taa acc tct	gct gaa ctg Ala Glu Leu 55	ggc tct anc ac Gly Ser Xaa Th	c tcc cyg 557 r Ser Xaa 60
gga aaa mta gca Gly Lys Xaa Ala 65	a tee age tot	gtg gcc tta Val Ala Leu 70	tcc gcg cat gt Ser Ala His Va 75	g cgg agg 605 l Arg Arg
ccc tgc gca gcc Pro Cys Ala Ala	c tgc agc agc a Cys Ser Ser	tac agc act Tyr Ser Thr	aag aga agc cc Lys Arg Ser Pr	c 647 o
80 tgagaaaaac ctc	tagaaaa acaaa	85 laaaaa aaaacc		684

<210> 306

<211> 693

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 262..471

<221> sig_peptide

<222> 262..306

<223> Von Heijne matrix
 score 3.5
 seq LCFLLPHHRLQEA/RQ

PCT/IB98/02122

<221> polyA_signal	•
<221> polyA_site <222> 682693	
<pre>&lt;400&gt; 306 atttcgcggc gctcgcbgma cyhsgwtgtt cagcaccttc ggtccggttg aggttgtcaa gtcggmccaa acaggttgtt tctctgcagt ttccaacatg gcagggmsgt ttaatagaca tggataagaa gtccactcac agaaatcctg aagatgccag ggctggcaaa tatgaaggta aacacaaacg aaagaaaaga agaaagcaaa accaaaacca gcaccgatcc cgacatagat cagtgacgtc ttttcttca g atg atc cta tgt ttc ctt ctt cct cat cat</pre>	60 120 180 240 291
cgt ctt cag gaa gcc aga cag att caa gta ttg aag atg ctt cca agg Arg Leu Gln Glu Ala Arg Gln Ile Gln Val Leu Lys Met Leu Pro Arg -5 1 5 10	339
gaa aaa tta aga aga aga gaa gag aga aaa caa ata aat ggg aaa aaa Glu Lys Leu Arg Arg Arg Glu Glu Arg Lys Gln Ile Asn Gly Lys Lys 15 20 25	387
raa agg aca aaa tat gaa aca cca aga aaa rga raa gga aaa aaa gga Xaa Arg Thr Lys Tyr Glu Thr Pro Arg Lys Xaa Xaa Gly Lys Lys Gly 30 35 40	435
gga aac mac cmc wtw tkt cmc ctt tcc aar agg gac tgaaactggg Gly Asn Xaa Xaa Xaa Xaa Leu Ser Lys Arg Asp 45 50 55	481
ctgacccttt tgatttccaa vctcascgtt ttggtgtaag gcggccaaar aaggatgcgg ascccagcac tgtgaagcct acaaaacat tgatgcgctg gcttggggat ttgaatttga acatctttca cactaagttc agactcatga aaccaatctt cagatgctct gtaaaccaca taataaagag tttggaaatt aaaaaaaaar aa	541 601 661 693
<210> 307 <211> 1656 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 741216	
<221> sig_peptide <222> 74172 <223> Von Heijne matrix score 5.80000019073486 seq XLCLGMALCPRQA/TR	
<221> polyA_signal	
<221> polyA_site <222> 16401652	
<pre>&lt;400&gt; 307 atctcttggc gtctcaacgt tcggatcagc agcttttttc cattctctct ctccacttct tcagtgagca gcc atg agt tgg act gtg cct gtt gtg cgg gcc agc cag</pre>	60 109
aga gtg agc tcg gtg gga gcg aat ktc cta tgc ctg ggg atg gcc ctg Arg Val Ser Ser Val Gly Ala Asn Xaa Leu Cys Leu Gly Met Ala Leu -20 -15 -10	157

																225
tgt	ccg	cgt	caa	gca	acg	cgc	atc	ccg	ctc	aac	ggc	acc	tgg	ctc	ttc	205
Cys	Pro	Arg	Gln	Ala	Thr	Arg	Ile	Pro		Asn	Gly	Thr	Trp	Leu	Pne	•
-5					1				5				- 4- 4-	10		253
acc	ccc	gtg	agc	aag	atg	gcg	act	gtg	aar	agt	gag	ctt	att	gag	cgt	253
Thr	Pro	Val	Ser	Lys	Met	Ala	Thr	Val	Lys	Ser	Glu	Leu	Ile	GIU	Arg	
			15					20					25			201
ttc	act	tcc	gar	aag	CCC	gtt	cat	cac	agt	aag	gtc	tcc	atc	ata	gga	301
Phe	Thr	Ser	Glu	Lys	Pro	Val	His	His	Ser	Lys	Val	ser	Ile	Ile	GIĀ	
		3.0					35					40				240
act	gga	tcg	gtg	ggc	atg	gcc	tgċ	gct	atc	agc	atc	tta	tta	aaa -	ggc	349
Thr	Gly	Ser	Val	Gly	Met	Ala	Cys	Ala	Ile	Ser	Ile	Leu	Leu	Lys	GIĀ	
	45					50					55					3.07
ttg	agt	gat	gaa	ctt	gcc	ctt	gtg	gat	ctt	gat	gaa	rac	aaa	ctg	aag	3 9 7
Leu	Ser	Asp	Ğlu	Leu	Ala	Leu	Val	Asp	Leu		Glu	Xaa	Lys	Leu	гув	
60					65					70					75	4.45
ggt	gag	acr	atg	gat	ctt	caa	cat	ggc	agc	cct	ttc	acg	aaa	atg	cca	445
Gly	Glu	Thr	Met	Asp	Leu	Gln	His	Gly	Ser	Pro	Phe	Thr	Lys	Met	Pro	
				80					85					90		400
aat	att	gtt	tgt	agc	aaa	rat	tac	ttt	gtc	aca	gca	aac	tcc	aac	cta	493
Asn	Ile	Val	Cys	Ser	Lys	Xaa	Tyr	Phe	Val	Thr	Ala	Asn	Ser	Asn	Leu	
			95					100					105			
gtg	att	atc	aca	gca	ggt	gca	cgc	caa	raa	aag	gga	gaa	acg	cgc	ctt	541
Val	Ile	Ile	Thr	Ala	Gly	Ala	Arg	Gln	Xaa	Lys	Gly	Glu	Thr	Arg	Leu	
		110					115					120				<b>500</b>
aat	tta	stc	cag	cga	aat	gtg	gcc	atc	ttc	aag	tta	atg	att	tcc	agt	589
Asn	Leu	Xaa	Gln	Arg	Asn	Val	Ala	Ile	Phe	Lys	Leu	Met	Ile	Ser	Ser	
	125					130					135					
att	gtc	cag	tac	agc	ccc	cac	tgc	aaa	ctg	att	att	gtt	tcc	aat	cca	637
Ile	Val	Gln	Tyr	Ser	Pro	His	Cys	Lys	Leu	Ile	Ile	Val	Ser	Asn	Pro	
140					145					150					722	
gtg	gat	atc	tta	act	tat	gta	gct	tgg	aag	ttg	agt	gca	ttt	ccc	aaa	685
Val	Asp	Ile	Leu	Thr	Tyr	Val	Ala	Trp	Lys	Leu	Ser	Ala	Phe	Pro	ьys	*
	_			160					165					170		
aac	cgt	att	att	gga	agc	ggc	tgt	aat	ctg	ata	mhg	gct	cgt	ttt	cgt	733
Asn	Arg	Ile	Ile	Gly	Ser	Gly	Cys	Asn	Leu	Ile	Xaa	Ala	Arg	Phe	Arg	
			175					180					185			
ttc	ttg	att	gga	caa	aag	ctt	ggt	atc	cat	tct	gaa	agc	tgc	cat	gga	781
Phe	Leu	Ile	Gly	Gln	Lys	Leu	Gly	Ile	His	Ser	Glu	Ser	Cys	His	Gly	
		190					195					200				
tgg	atc	ctc	gga	gag	cat	gga	gac	tca	agt	gtt	cct	gtg	tgg	agt	gga	829
Trp	Ile	Leu	Gly	Glu	His	Gly	Asp	Ser	Ser	Val	Pro	Val	Trp	Ser	Gly	
	205					210					215					
gtg	aac	ata	gct	ggt	gto	cct	ttg	aag	gat	ctg	aac	tct	gat	ata	gga	877
Val	Asn	Ile	Ala	Gly	Val	Pro	Leu	Lys	Asp	Leu	Asn	Ser	Asp	Ile	GIY	
220	)				225	,				230	)				235	
act	gat	aaa	gat	cct	gag	caa	. tgg	aaa	aat	gto	cac	aaa	gaa	gtg	act	925
Thr	Asp	Lys	Asp	Pro	Glu	Gln	Trp	Lys	Asn	Val	. His	Lys	Glu	. Val	rnr	
				240	)				245	,				250	)	
gca	act	gco	tat	gag	att	att	aaa	atg	aaa	ggt	: tat	act	tct	tgg	gcc	973
Āla	Thr	Ala	Tyr	Glu	ı Ile	: Ile	Lys	Met	. Lys	Gly	, Tyr	Thr	Ser	Trp	Ala	
			255	;				260	)				265	•		
att	ggc	cta	tct	gtg	gco	gat	: tta	ı aca	gaa	ı agt	att	ttg	aag	, aat	ctt	1021
Ile	Gly	Let	ser	Val	Ala	Asp	Let	Thr	Glu	. Ser	: Ile	Let	ı Lys	Asn	Leu	
		270	)				275	5				280	)			
agg	aga	ata	a cat	. cca	gtt	tcc	acc	: ata	act	aaq	999	: ctc	: tat	: gga	ata	1069
Arc	Arc	, Ile	e His	Pro	va]	Ser	Thr	: Ile	Thr	Lys	: Gly	/ Let	Tyr	: Gly	/ Ile	
-	285	;				290	)				295	5				
rat	gaa	gaa	a gta	tto	cto	agt	: att	cct	: tgt	ato	cts	g gga	gag	g aac	ggt	1117
Xaa	a Glu	ı Ğlı	ı Val	Phe	e Lei	ı Ser	: Ile	e Pro	Сув	: Ile	e Lev	ı Gly	/ Glu	1 Asr	J GTA	
300	)			4	305	5				310	)				315	
ati	t acc	: aa	ctt	: ata	aaa	g ata	aaa	g cto	gaco	cct	gaa	gaa	gaç	a áco	cat	1165
Ile	e Thi	: As	n Lev	ı Ile	E Ly	s Ile	E Lys	s Lev	ı Thi	e Pro	o Gli	ı Glu	ı Glu	ı Ala	a His	

220	
320  Ctg aaa aaa agt gca aaa aca ctc tgg gaa att cag aat aag ctt aag  Leu Lys Lys Ser Ala Lys Thr Leu Trp Glu Ile Gln Asn Lys Leu Lys	1213
335 340 345 ctt taaagttgcc taaaactacc attccgaaat tattgaagag atcatagata	1266
Len	1326
caggattata taacgaaatt ttgaataaac ttgaattcct aaaagatgga aacaggaaag	1386
taggtagagt gattttccta tttatttagt cctccagctc ttttattgag catccacgtg	1446
ctggacgata cttatttaca attcckaagt atttttggta cctctgatgt agcagcactt gccatgttat atatatgtag ttgrmatttg gttcccaaaa agtaggatgt aggtatttat	1506
tgtgttctag aaattccgac tcttttcatt agatatatgc tatttctttc attcttgctg	1566
gtttatacct atgttcattt atatgctgta aaaaagtagt agcttcttct acaatgtaaa	1626
aataaatgta catacaaaaa aaaaaamcmc	1656
<210> 308	
<211> 517	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 48164	
<221> sig_peptide	
<222> 4889	
<223> Von Heijne matrix	
score 4	
seq YYMVCLFFRLIFS/EH	
<221> polyA_signal	
<222> 482487	
\2227 \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \	•
<221> polyA_site	
<222> 505517	
<400> 308	56
aggagatagc ctcgtagaaa tgacaaccac aatgttaata ctaacat atg tat tac Met Tyr Tyr	
atg gtt tgt ttg ttc ttt cgc tta ata ttt tca gag cac cta ctt att	104
Met Val Cys Leu Phe Phe Arg Leu Ile Phe Ser Glu His Leu Pro Ile	
-10 -5	
ata ggc act gtc act tct cac aaa act ggg aca cta act gtt tat cca	152
Ile Gly Thr Val Thr Ser His Lys Thr Gly Thr Leu Thr Val Tyr Pro	
10 15 20	
aca tot got ggo taaataaaga catgatotto accttttggg attgttaatt	204
Thr Ser Ala Gly	
25	264
taaaatggtt ccataagagc aatgcaaaga cagagatatt tggcagcact gcagctggtg	324
atttatatgg ctcttcacaa ggtgttattt tggggtatca aggtatggat gcttaaatca gctgcaggaa gtaagaaaga agaaaaaagg agtgataaag ataaaaaaaa	384
gccgcaggaa gtaagaaaga agaaaaaagg agcgacaaag acaaaaattcc gcccttccac caaaacccat taatttccat atcatcatct gcataarara gaaaattcc	444
acwtgaccag gttactgcaa ggatktkaat tttgaatatt aaaatattat mcmcaattgg	504
aaaaaaaaaa aaa	517

<211> 405

<212> DNA

<213> Homo sapiens

<220> <221> CDS <222> 185334	
<pre>&lt;221&gt; sig_peptide &lt;222&gt; 185295 &lt;223&gt; Von Heijne matrix</pre>	
<221> polyA_signal	
<221> polyA_site   <222> 392405	
<pre>&lt;400&gt; 309 atcaccttct tctccatcct tstctgggcc agtccccarc ccagtccctc tcctgacctg cccagcccaa gtcagccttc agcacgcgct tttctgcaca cagatattcc aggcctacct ggcattccag gacctccgma atgatgctcc agtcccttac aagcgcttcc tggatgaggg tggc atg gtg ctg acc acc ctc ccc ttg ccc tct gcc aac agc cct gtg     Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val</pre>	60 120 180 229
aac atg ccc acc act ggc ccc aac agc ctg agt tat gct agc tct gcc Asn Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala -20 -15 -10	277
ctg tcc ccc tgt ctg acc gct cca aag tcc ccc cga ctt gct atg atg Leu Ser Pro Cys Leu Thr Ala Pro Lys Ser Pro Arg Leu Ala Met Met -5 1 5 10	325
cct gac aac taaatatcct tatccaaatc aataaarwra raatcctccc Pro Asp Asn	374
<pre>tccaraaggg tttctaaaaa caaaaaaaa a  &lt;210&gt; 310 &lt;211&gt; 1087 &lt;212&gt; DNA &lt;213&gt; Homo sapiens</pre>	
<220>	
<221> CDS <222> 195347	
<221> sig_peptide <222> 195272 <223> Von Heijne matrix score 7.09999990463257 seq LASLQWSLTLAWC/GS	
<221> polyA_signal	
<221> polyA_site <222> 10711082	
<400> 310 aaagtgtaga acacggacct ctgagttatg ctcttgagag gtgccaaagc tgggctgttt acctacctta tccacagagc tctgaaagtc aagccagaaa ggaaggattc caaattcttg gaattttatc tagaaaagaa gactaagcag cttttgttct tctgtgaccc agttgctggc caaattctagcagagagagagagagagagagagagagaga	60 120 180 230

Met Thr Pro Trp Cys Leu Ala Cys Leu Gly Arg Arg	
-25 -20 -15  cct ctc gct tct ttg cag tgg agc ctg aca ctg gcg tgg tgt ggc tcc  Pro Leu Ala Ser Leu Gln Trp Ser Leu Thr Leu Ala Trp Cys Gly Ser	278
-10 -5 ¹	226
ggc agc cac tgg aca gag aga cca akt cag akt tca ccg tgg akt tct Gly Ser His Trp Thr Glu Arg Pro Xaa Gln Xaa Ser Pro Trp Xaa Ser 5 10 15	326
ctg tca gcg acc acc agg ggg tgatcacacg gaaggtgaac atccaggtcg	377
Leu Ser Ala Thr Thr Arg Gly 20 25	
gggatgtgaa tgacaacgcg cccacatttc acaatcagcc ctacagcgtc cgcatccctg	437
araatacacc agtggggacg cccatcttca tcgtgaatgc cacagacccc gacttggggg	497 557
cagggggcag cgtcctctac tccttccagc cccctccca attcttcgcc attgacagcg cccgcggtat cktcacagtg atccgggagc tggactacga taccacrcmg gcctaccagc	617
towoggtowa ogcoacagat caagacaara coaggootot gtocacostg gocaacttgg	677
ccatcatcat cacagatote caogacatog accepatett catcaaceto cettacagea	737
ccaacatcta cgagcattot cotocqqqoa cgaeggtgeg cateateace gecatagaee	797
aggataaagg acgtccccgg ggcattggct acaccatcgt ttcagggcat ctgtgtttac	857 917
aagaacccaa gatctctcag gagctcagga aaaggggctt gctgtgaggc tcagggttcc catggacatt ctgagctgac cctcctcagc attggatctc ctggctcagg aactaggaac	977
gaagettgga tgttttetee ttteetacag catetgtatt cattteetat agttgecata	1037
ataaaatgcc actaacttag tggcttaaaa accaaaaaaa aaaaaccctt	1087
<210> 311	
<211> 916	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS <222> 90815	
<221> sig_peptide	
<222> 90179 <223> Von Heijne matrix	
score 13.1999998092651	
seq LLLLSTLVIPSAA/AP	
<221> polyA signal	
<222> 883888	
<221> polyA site	
<222> 905916	
<400> 311	
aaaacaqtac qtqqqqqqc qqaatccggg agtccggtga cccgggctgt ggtctagcat	60
aaaggcggag ccagaagaag gggcggggt atg gga gaa gcc tcc cca cct gcc	113
Met Gly Glu Ala Ser Pro Pro Ala	
-30 -25 ccc gca agg cgg cat ctg ctg gtc ctg ctg ctc ctc tct acc ctg	161
Pro Ala Arg Arg His Leu Leu Val Leu Leu Leu Leu Ser Thr Leu	
-20 -15 -10	
gtg atc ccc tcc gct gca gct cct atc cat gat gct gac gcc caa gag	209
Val Ile Pro Ser Ala Ala Ala Pro Ile His Asp Ala Asp Ala Gln Glu	
age tee ttg ggt etc aca gge etc cag age etc caa gge tte age	257
Ser Ser Leu Gly Leu Thr Gly Leu Gln Ser Leu Leu Gln Gly Phe Ser	
15 20 25	205
cga ctt ttc ctg aaa ggt aac ctg ctt cgg ggc ata gac agc tta ttc	305

_			30					35					40	Leu			
+ < +	acc	CCC	ato	gac	ttc	caa	aac	ctc	cct	aaa	aac	tac	cac	aaa	gag	3	353
-	33-	200	Mak	3	Dha	722	631	Len	Pro	Gly	Aen	ጥህጕ	His	Lys	Glu		
ser	Ата		Met	Asp	Pne	Arg		neu	FIO	Cly	7311	- 7 -		-1-			
		45					50					55					
gag	aac	cag	gag	cac	cag	ctg	999	aac	aac	acc	ctc	tcc	agc	cac	CEC	-	401
Glu	Asn	Gln	Glu	His	Gln	Leu	Gly	Asn	Asn	Thr	Leu	Ser	Ser	His	Leu		
	60					65	-				70						
		~~~	224	3 t G	200		aac	aad	aca	gga	gag	ata	cta	atc	tcc	4	449
cag	all	gac	aay	acg	mb	3	700	Tura	Thr	25~	G111	Val	T.ell	Tle	Ser		
	IIe	Asp	гÀв	Met		Asp	ASII	пур	1111	GIY	GIU	Val	200	Ile	90		
75					80					85							407
gag	aat	gtg	gtg	gca	tcc	att	caa	cca	vcg	gag	999	anc	ttc	gag	ggt	•	497
Glu	Asn	Val	Val	Ala	Ser	Ile	Gln	Pro	Xaa	Glu	Gly	Xaa	Phe	Glu	Gly		
				95					100					105			
~a+	++~	220	ath		agg	ato	gag	gar	aaq	gag	acc	cta	qta	ccc	mtc		545
gat	T	aag	9011	D~-	722	Mat	Glu	Glu	Lve	Glu	Δla	T.eu	Val	Pro	Xaa		
Asp	Leu			PIO	Arg	Mec	GIU		נעם	Oru			120				
			,110					115									593
car	aag	gcc	acg	gac	agc	ttc	cac	aca	gaa	ctc	cat	ccc	cgg	gtg	gcc	•	553
Gln	Lys	Ala	Thr	Asp	Ser	Phe	His	Thr	Glu	Leu	His	Pro	Arg	Val	Ala		
	-	125					130					135					
ttc	t.aa	atc	att	aaq	cta	cca	cqq	cgg	agg	tcc	cac	cag	gat	gcc	ctg	1	641
Dhe	Trn	Tle	Tle	Lvs	Len	Pro	Ara	Arg	Ara	Ser	His	Gln	Asp	Ala	Leu		
FIIC	140			-1-		145					150		-				
							~~~	~~~		a . a		ata	cac	acc	atc		689
gag	ggc	ggc	cac	tgg	CEC	anc	gar	aag	cya	Cac	200	t	01-	gcc	Tlo		• • •
Glu	Gly	Gly	His	Trp	Leu	Xaa	Glu	Lys	Arg	HIS	Arg	ьeu	GIN	Ala	116		
155					160					165					170		
caa	gat	qqa	ctc	cqc	aag	ggg	acc	cac	aag	gac	rtc	cta	daa	rag	ggg		737
Ara	Asn	Glv	Len	Ara	Lvs	Glv	Thr	His	Lys	Asp	Xaa	Leu	Xaa	Xaa	Gly		
5		<b>-</b> -3		175	-2-				180	-				1.85			
	~~~	200	+		C 2 C	tcc	200	cta		CCC	cga	aar	amm	cac	tta		785
acc	gar	age	0	2	TT:	000	723	IOU	200	220	720	Tare	Yaa	His	Leu		
Thr	Glu	ser		Ser	HIS	Ser	Arg		Ser	PIO	AL 9	Буз	200				
			190					195									835
											gggt	999	gacc	aaaa	ar		033
Leu	Tyr	Ile	Leu	Xaa	Pro	Ser	Arg	Gln	Leu								
	_	205					210										
mac	ctac	cta	tage	cccc	at c	arac	cctq	c cc	caag	cacc	ata	tgga	aat	aaag	ttcttt	:	895
				aaaa					_								916
	acat	cea	uaaa	uaua	~ u												

<211> 583

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 52..513

<221> sig_peptide

<222> 52..231 <223> Von Heijne matrix score 4 seq LVRRTLLVAALRA/WM

<221> polyA_signal

<222> 553..558

<221> polyA_site <222> 572..583

<400> 312

aaggaaacag caaccagagg gagatgatca cctgaaccac tgctccaaac c atg ggc Met Gly -60	57
agt aaa tgc tgt aaa ggt ggt cca gat gaa gat gca gta gaa aga cag Ser Lys Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu Arg Gln -55 -50 -45	105
agg cgg cag aag ttg ctt ctt gca caa ctg cat cac aga aaa agg gtg Arg Arg Gln Lys Leu Leu Ala Gln Leu His His Arg Lys Arg Val -40 -35 -30	153
aar gca gct ggg cag atc cag gcc tgg tgg cgt ggg gtc ctg gtg cgc Lys Ala Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu Val Arg -25 -20 -15	201
agg acc ctg ctg gtt gct gcc ctc agg gcc tgg atg att cag tgc tgg Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln Cys Trp -10 -5 1 5	249
tgg agg acg ttg gtg cag aga cgg atc cgt cag cgg cgg cag gcc ctg Trp Arg Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln Ala Leu 10 15 20	297
ttr ggg gtc tac gtc atc cag gag cag gcg gcg gtc aag ctc cag tcc Leu Gly Val Tyr Val Ile Gln Glu Gln Ala Ala Val Lys Leu Gln Ser	345
tgc atc cgc atg tgg cag tgc cgg caa tgt tac cgc caa atg tgc aat Cys Ile Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met Cys Asn 40 45 50	393
gct ctc tgc ttg ttc cag gtc cca aaa agc agc ctt gcc ttc caa act Ala Leu Cys Leu Phe Gln Val Pro Lys Ser Ser Leu Ala Phe Gln Thr 55 60 65 70	441
gat ggc ttt tta cag gtc caa tat gca atc cct tca aag cag cca gag Asp Gly Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln Pro Glu 75	489
ttc cac att gaa atc cta tca atc tgaaaggcct ggggcatgga gaacaggctg Phe His Ile Glu Ile Leu Ser Ile 90	543
cactacccta ataaatgtct gaccaggtaa aaaaaaaaaa	583

<211> 697

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 172..438

<221> sig_peptide

<222> 172..354

<223> Von Heijne matrix score 4.69999980926514 seq LLPCNLHCSWLHS/SP

<221> polyA_signal

<222> 682..687

<221> polyA_site

<222> 685..697

<400> 313

60 agattggctg ggcagatggg ctgactggct gggcagatgg gtgggtgagt tccctctccc cagagocato ggocaggtac caaagotcag otgtatggat toccaacagg aggacotgog 120 cttccctggg acccattgtt gtactggatt aacaagcgac ggcgctacgg c atg aat 177

Met Asn	
-60	225
gca gcc atc aac acg ggc cct gcc cct gct gtc acc aag act gag act Ala Ala Ile Asn Thr Gly Pro Ala Pro Ala Val Thr Lys Thr Glu Thr	223
-55 -50 -45	
gag gtc cag aat cca gat gtt ctg tgg gat ttg gac atc ccc gaa gcc	273
Glu Val Gln Asn Pro Asp Val Leu Trp Asp Leu Asp Ile Pro Glu Ala	
agg agc cat gct gac caa gac agc aac ccc aag gcg gaa gcc ctg ctc	321
Arg Ser His Ala Asp Gln Asp Ser Asn Pro Lys Ala Glu Ala Leu Leu	
-25 -20 -15	360
ccc tgc aac ctg cac tgc agc tgg ctc cac agc agc ccc agg cca gat	369
Pro Cys Asn Leu His Cys Ser Trp Leu His Ser Ser Pro Arg Pro Asp -10 -5 1 5	
ccc cat tcc cac ttc cca tct ktc agg agg tgc cct ttg ccc cac cct	417
Pro His Ser His Phe Pro Ser Xaa Arg Arg Cys Pro Leu Pro His Pro	
10 15 20	468
tgt gca acc tac ccc ccs kgc tgaaccactc tgtctcctat cctttggcca	400
Cys Ala Thr Tyr Pro Pro Xaa 25	
cotgeoetga aaggaatget otottocatt cootcotgaa totggoocag gaagaccata	528
getteaatgy caageetttt cetteaaaac tgtageetee teteaetgaa ggtgggaget	588
gcaggaatca ggtgcagagt aggaaatgga actaacctca ggaaggtggt attgacagag	648
gtcaggaccc acctggatgt catgctatga aacattaaaa gaaaaaaaa	697
<210> 314	
<211> 803	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 148366	
<221> sig_peptide	
<222> 148225	
<pre><223> Von Heijne matrix score 5.5</pre>	
seg LFTLLFLIMLVLK/LD	
504 II. 1111 III	
<221> polyA_signal	
<222> 770775	
<221> polyA_site <222> 792803	
2227 192003	
<400> 314	
aaatgggggg aaaagggcgg aaaaggacaa ggatccaaac tggcgaattt gctgatcttc	60
gegteeetet eegettteeg geeggeageg etgeeagggt atattteett tttteegate	120
ctgcaacage ctetttaaac tgtttaa atg aga atg tee ttg get cag aga gta	174
Met Arg Met Ser Leu Ala Gln Arg Val	
cta ctc acc tgg ctt ttc aca cta ctc ttc ttg atc atg ttg gtg ttg	222
Leu Leu Thr Trp Leu Phe Thr Leu Leu Phe Leu Ile Met Leu Val Leu	
-15 -10 -5	
aaa ctg gat gag aaa gca cct tgg aac tgg ttc ctc ata ttc att cca	270
Lys Leu Asp Glu Lys Ala Pro Trp Asn Trp Phe Leu Ile Phe Ile Pro	
1 5500 15 15 15 15 15 15 15 15 15 15 15 15 15	318
gtc tgg ata ttt gat act atc ctt ctt gtc ctg ctg att gtg aaa atg Val Trp Ile Phe Asp Thr Ile Leu Leu Val Leu Leu Ile Val Lys Met	3.0
the the tre and the tre and and the can are the tree at	

25 30	
gct ggg cgg tgt aag tct ggc ttt gac ctc gac atg gat cac aca ata Ala Gly Arg Cys Lys Ser Gly Phe Asp Leu Asp Met Asp His Thr Ile	366
રુદ્	426
taaaaaaaaa aacctggtac ctcattgcac tgtkacttaa attasccttc tgcctcgcac	486
tetatatatata actggaacag tttactacca tgaatctate etatgtette atteetttat gggeettget ggetgggget ttaacagaac teggatataa tgtetttttt gtgaaagact	546
gacttetaag tacatcatet cetttetatt getgtteaac aagttaceat taaagtgtte	606
tgaatctotc aagcttcaag aataccagag aactgaggga aaataccaaa tgtagtttta	666
tactacttcc ataaaacagg attggtgaat cacggacttc tagtcaacct acagcttaat	726
tattcagcat ttgagttatt gaaatcctta ttatctctat gtaaataaag tttgttttgg	786
acctcaaaaa aaaaaaa	803
•	
<210> 315	
<211> 823	
<212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS	
<222> 175336	
<221> sig peptide	
<222> 175276	
<223> Von Heijne matrix	
score 3.7000004768372	
seq SVLNVGHLLFSSA/CS	
<221> polyA site	
<222> 812823	
<400> 315	60
aaggegegeg egaceggegg etetttggeg eggattaggg ggteteggeg agggagteat	120
caagetttgg tgtatgtgtt ggeeggttet gaagtettga agaagetetg etgaggaaga ecaaageage actegttgee aattagggaa tggaeegttt gggtteettt agea atg	177
Met	
ate cet etg ata age cae ett gee gag get get eet eet aee tea tgg	225
Ile Pro Leu Ile Ser His Leu Ala Glu Ala Ala Pro Pro Thr Ser Trp	
-30 -25 -20	273
ago ott ata toa agt gtg etg aat gtg gge cac ete ett tit toe tet	273
Ser Leu Ile Ser Ser Val Leu Asn Val Gly His Leu Leu Phe Ser Ser	
-15 -10 -5 gct tgc agt gtt tca ctc gag gct ttg agt aca aga aac atc aaa gcg	321
Ala Cys Ser Val Ser Leu Glu Ala Leu Ser Thr Arg Asn Ile Lys Ala	
1 5 10 15	
atc ata ctt atg aaa taatggette agatttteet gteettgate eeagetggae	376
Ile Ile Leu Met Lys	
20	436
tgctcaagaa raaatggccc ttttagaasc tgtgatggac tgtggctttg gaaattggca ggatgtagcc aatcaaatgt gcaccaarac caaggaggag tgtgagaagc actatatgaa	496
gcatttcatc aataacccyc tgtttgcatc trscctgctg aacctgaaac aascagrgga	556
agcaaaaact gctgacacag ccattccatt tcactctaca ratgaccctc cccgacckac	616
ctttgactcc ttgctttctc gggacatggc cgggtacwtg ccmgctcgag cagatttcat	676
tgaggaattt gacaattatg cagaatggga cttgagagac attgattttg ttgaagatga	736 796
ctcggacatt ttacatgctc tgaagatggc tgtggtagat atctatcatt ccaggttaaa	823
ggagagacaa agacgaaaaa aaaaaaa	

```
<211> 823
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 191..553
<221> sig_peptide
<222> 191..304
<223> Von Heijne matrix
     score 5.69999980926514
      seq LAFLSCLAFLVLD/TQ
<221> polyA_signal
<222> 766..771
<221> polyA_site
<222> 804..817
<400> 316
                                                                      60
aactotgoag ggcotocaag gccaggotto agggotggga otcagtootg aggcactggg
                                                                      120
gagecatgag gggctgtggc agggagggc agggtgtgga aagaeteeee tggggecatg
gtggagatgt gctgaggtct tctccctgat cgtcttctcc tccctgctga ccgacggcta
                                                                      180
ccagaackag atg gag tot ccg cag ctc cac tgc att ctc aac agc aac
                                                                      229
          Met Glu Ser Pro Gln Leu His Cys Ile Leu Asn Ser Asn
                                           -30
                       -35
                                                                      277
age gtg gee tge age ttt gee gtg gga gee gge tte etg gee tte ete
Ser Val Ala Cys Ser Phe Ala Val Gly Ala Gly Phe Leu Ala Phe Leu
                                         -15
                                                             -10
                    -20
-25
age tgc ctg gcc ttc ctc gtc ctg gac aca cag gag acc cgc att gcc
                                                                      325
Ser Cys Leu Ala Phe Leu Val Leu Asp Thr Gln Glu Thr Arg Ile Ala
                -5
ggc acc cgc ttc aag aca gcc ttc cag ctc ctg gac ttc atc ctg gct
                                                                      373
Gly Thr Arg Phe Lys Thr Ala Phe Gln Leu Leu Asp Phe Ile Leu Ala
        10
                            15
gtt ctc tgg gca gtt gtc tgg ttc atg ggt ttc tgc ttc ctg gcc aac
                                                                      421
Val Leu Trp Ala Val Val Trp Phe Met Gly Phe Cys Phe Leu Ala Asn
                        30
                                                                      469
caa tgg cag cat tcg ccg ccc aaa gar kkc ctc ctg ggg agc agc agt
Gln Trp Gln His Ser Pro Pro Lys Glu Xaa Leu Leu Gly Ser Ser Ser
40
                                         50
ged cag gea ged ate ggc stt cac ett ett etc eat eet tgt etg gat
                                                                      517
Ala Gln Ala Ala Ile Gly Xaa His Leu Leu His Pro Cys Leu Asp
                60
                                    65
                                                                      563
att cca rgc cta cct ggc akk cca gga cct ccg aaa tgatgctcca
Ile Pro Xaa Leu Pro Gly Xaa Pro Gly Pro Pro Lys
            75
gtcccttacm arcgcttcct ggatgaaggt ggcatggtgs kkaacaccct ccccttgccc
                                                                      623
                                                                      683
totgccaaca gootgtgaac atgcccacca otggccccaa cagcotgagt tatgctagct
ctgccctgtc cccctgtctg accgctcmaa agtccccccg gcttgctatg atgcctgaca
                                                                      743
                                                                      803
actaaatatc cttatccaaa tcaataaaga gagaatcctc cctccagaag ggtttctaaa
                                                                      823
aacaaaaaa aaaahncctt
```

<211> 1112

<212> DNA

<213> Homo sapiens

<221> CDS <222> 106..603 <221> sig_peptide <222> 106..216 <223> Von Heijne matrix score 4.30000019073486 seq LWEKLTLLSPGIA/VT <221> polyA_site

<221> polyA_site <222> 1102..1112

<400> 317 60 agegattgcg aatcctccgc tgaggtgatt tggatatccc tagaacgttg agggcacgag 117 tegggteetg agaccaggte etcagecage agagecaegt teett atg age ace gtg Met Ser Thr Val ggt tta ttt cat ttt cct aca cca ctg acc cga ata tgc ccg gcg cca 165 Gly Leu Phe His Phe Pro Thr Pro Leu Thr Arg Ile Cys Pro Ala Pro -25 tgg gga ctc cgg ctt tgg gag aag ctg acg ttg tta tcc cca gga ata 213 Trp Gly Leu Arg Leu Trp Glu Lys Leu Thr Leu Leu Ser Pro Gly Ile -10 -15 261 get gte act eeg gte cag atg gea gge aag aag gae tae eet gea etg Ala Val Thr Pro Val Gln Met Ala Gly Lys Lys Asp Tyr Pro Ala Leu 10 309 ctt tcc ttg gat gag aat gaa ctc gaa gag cag ttt gtg aaa gga cac Leu Ser Leu Asp Glu Asn Glu Leu Glu Glu Gln Phe Val Lys Gly His 20 25 ggt cca ggg ggc cag gca acc aac aaa acc agc aac tgc gtg gtg ctg 357 Gly Pro Gly Gly Gln Ala Thr Asn Lys Thr Ser Asn Cys Val Val Leu 40 405 aar mac atc ccc tca ggc atc gtt gta aag tgc cat cag aca aga tca Lys Xaa Ile Pro Ser Gly Ile Val Val Lys Cys His Gln Thr Arg Ser 55 gtt gat cag aac aga aag cta gct cgg aaa atc cta caa gag aaa gta Val Asp Gln Asn Arg Lys Leu Ala Arg Lys Ile Leu Gln Glu Lys Val 70 501 rat gtt ttc tac aat ggt gaa aac agt cct gtt cac aaa gaa aaa cga Xaa Val Phe Tyr Asn Gly Glu Asn Ser Pro Val His Lys Glu Lys Arg 90 85 80 549 gaa gcg gcg aag aaa aaa car gaa agg aaa aaa aga gca aag gaa acc Glu Ala Ala Lys Lys Gln Glu Arg Lys Lys Arg Ala Lys Glu Thr 105 100 597 ctg gaa aaa aag aas ctm ctt aaa raa ctg tgg gag tca agt aaa aag Leu Glu Lys Lys Xaa Leu Leu Lys Xaa Leu Trp Glu Ser Ser Lys Lys 120 653 gtc cac tgagaaaaga attagagatt ccaactgaca gaatctgcca gaagctccca Val His 713 gggaataatg gtggcgagtt ccatcaccag cattattata gtgcttcaaa agaaatattt 773 ttgatgaact taaaagacaa caaatttatt taaatggtgc actaaactgt agtgaacaga gacatgcacg attcaagaat aaaactcggc cgggcacggt ggacggtgcc tcacatctgt 833 aatcccagca ctttgggagg ccgaggcggg cggatcactt gaggtcagga gtttgagacc 893 agcctggcca acatggtgaa accccgtctc tactaaaaat acaaaaaatt agccaggcat 953 ggtggcgggc acctgtaatc ccagctactc gggaggccga ggcaggagaa ttgcgtgaac 1013 1073 ctgggaggcg gaggttgcag tgagctgaga tcgcgccact gcactcaagc ctgggcaaca 1112 cctgggtgac agagcaagac cccatcycaa aaaaaaaaa

WO 99/31236

<212> DNA <213> Homo sapiens <220> <221> CDS <222> 47..586 <221> sig_peptide <222> 47..124 <223> Von Heijne matrix score 6.30000019073486 seq GVGLVTLLGLAVG/SY <221> polyA_signal <222> 1583..1588 <221> polyA_site <222> 1614..1623 <400> 318 55 agggatetgt eggettgtea ggtggtggag gaaaaggege teegte atg ggg ate Met Gly Ile cag acg age eec gte etg etg gee tee etg ggg gtg ggg etg gte act 103 Gln Thr Ser Pro Val Leu Leu Ala Ser Leu Gly Val Gly Leu Val Thr -15 ctg ctc ggc ctg gct gtg ggc tcc tac ttg gtt cgg agg tcc cgc cgg 151 Leu Leu Gly Leu Ala Val Gly Ser Tyr Leu Val Arg Arg Ser Arg Arg cct cag gtc act ctc ctg gac ccc aat gaa aag tac ctg cta cga ctg Pro Gln Val Thr Leu Leu Asp Pro Asn Glu Lys Tyr Leu Leu Arg Leu 20 10 cta gac aag acg act gtg agc cac aac acc aag agg ttc cgc ttt gcc Leu Asp Lys Thr Thr Val Ser His Asn Thr Lys Arg Phe Arg Phe Ala 30 35 295 ctg ccc acc gcc cac cac act ctg ggg ctg cct gtg ggc aaa cat atc Leu Pro Thr Ala His His Thr Leu Gly Leu Pro Val Gly Lys His Ile 50 343 tac ctc tcc acm mga att gat ggc agc ctg gtc atc agg cca tac act Tyr Leu Ser Thr Arg Ile Asp Gly Ser Leu Val Ile Arg Pro Tyr Thr 65 cct gtc acc agt gat gag gat caa ggc tat gtg gat ctt gtc mtc aag 391 Pro Val Thr Ser Asp Glu Asp Gln Gly Tyr Val Asp Leu Val Xaa Lys 8.0 gtc tac ctg aag ggt gtg cac ccc aaa ttt cct gag gga ggg aar atg 439 Val Tyr Leu Lys Gly Val His Pro Lys Phe Pro Glu Gly Gly Lys Met 100 487 tct cak tac ctg gat asc ctg aaa gtt ggg gat btg gtg gaa ttt csg Ser Xaa Tyr Leu Asp Xaa Leu Lys Val Gly Asp Xaa Val Glu Phe Xaa 115 110 535 ggg cca agc ggg ttg ctc act tac act gga aaa ggg cat ttt aac att Gly Pro Ser Gly Leu Leu Thr Tyr Thr Gly Lys Gly His Phe Asn Ile 130 583 cag ccc aac aag aat ctc cac cag aac ccc gag tgg cga aga aac tgg Gln Pro Asn Lys Asn Leu His Gln Asn Pro Glu Trp Arg Arg Asn Trp gaa tgattgccgg cgggacagga atcaccccaa tgctacagct gatccgggcc 636 Glu atcctgaaag tccctgaaga tccaacccag tgctttctgc tttttgccaa ccagacagaa 696 756 aaggatatca tettgeggga ggaettagag gaactgeagg ecegetatee caategettt aagetetggt teactetgga teatececea aaagrttggg cetacageaa gggetttgtg 816 actgccgacw tgatccggga acacctgccc gctccagggg atgatgtgct ggtactgctt 876

tgtgggccmc ccccaatggt gca tcacaaaga tgcgattcac cta agttgttccc catcagtact caa ggtttttca gttrsatcka ga gtagccatgg aagagggcca aga tggcaacagg tccaggagag gca agcatgtacg cttggtccaa ga ttgtgtctgt gatgaaagga ac acctatgaga atggcaagaa aga tcttatggag atggcaagaa aga tatcatattt ctgtgtgtgt cda gataatcgaa aactgctgtt tg ngccctgtgt gataattgaaa aa	actgagea tectecaget ageactak aageettagr etggatagta geteagte acteettgga eagtetette ttggetag tecttgata agtetgtg caatgggtt gtgagtat aagttgagea gaggaaat gatttettea eteteage eeetgeeead tgggearg aaeeeetgge	ktcctktcct cagagtttca cctgcaggaa caatattcct tggcctccta aatctccccg catggagtaa gaaggaaggg gcatcttact ctcaccttct tacttaaact tcactgttca tagcatactt ccagaggtgg gatctcaaag gagtctgaaa gctagagga wacagctact	936 996 1056 1116 1176 1236 1296 1356 1416 1476 1536 1596 1623
<210> 319			
<211> 526 <212> DNA			
<213> Homo sapiens			
<220> <221> CDS			
<222> 99371			
<221> sig_peptide <222> 99290			
<pre><223> Von Heijne matrix score 3.7999999523</pre>	1628		
seq LFIVVCVICVTLN/			
<221> polyA_signal <222> 491496			
<221> polyA_site <222> 513524			
<400> 319		atatattat ttaggacatt	60
attggattag tagaattgct tt ttactttttt ctgttaacgc tt	accctagr aattagaa a	tg aca cca cgt att ctt let Thr Pro Arg Ile Leu	116
agc gaa gtc cag ttt tca	gca ttt tgt cct tat	tgg aca ata gca agg	164
Ser Glu Val Gln Phe Ser -55	-50	-45	
ata tta gaa cgt gtt ggt Ile Leu Glu Arg Val Gly	tcc gcg tgc ttc cgt Ser Ala Cys Phe Arg	ctt gag tta tgt gct Leu Glu Leu Cys Ala	212
-40 get att gte gga tat ttt	-35	-30	260
Ala Ile Val Gly Tyr Phe	Val Leu Asp Val Arg	Thr Phe Leu Phe Ile	
-25 gtg gta tgt gta att tgc	gtt act ttg aat ttt	cca cgt ttt tac ttt	308
Val Val Cys Val Ile Cys -10 -5	1	5	256
ctt tgt ctc tca tca ctt Leu Cys Leu Ser Ser Leu	acc gct ttt ggg acc Thr Ala Phe Gly Th	c ccc ccc atc ggg gtt c Pro Pro Ile Gly Val	356
10 cac att ccc tct ccc tar	15	20	411
His Ile Pro Ser Pro			
gcggtgaage tttcccattt t	atgtgcaga ttattttcag	g agggtatata gaattcaggc	471 526
agetgttteg ttgtageaca t	Laadaalat tttttttttttt		

<210> 320 <211> 989 <212> DNA <213> Homo sa	apiens					
<220> <221> CDS <222> 44814	4					
	2					
<221> polyA_s <222> 97898						
<400> 320 aaatgtgtac a	cgcccagct t	cctgcctgt 1	tactctccac	agt atg cga Met Arg	aga ata Arg Ile -20	55
tcc ctg act s	tct agc cct Ser Ser Pro -15	gtg cgc ct Val Arg L	tt ctt ttg eu Leu Leu -10	tdt ctg ctg Xaa Leu Leu	ttg cta Leu Leu -5	103
cta ata gcc : Leu Ile Ala :	tto gag atc	atg gtt g Met Val G	gt ggt cac ly Gly His	tct ctt tgc Ser Leu Cys 10	ttc aac Phe Asn	151
ttc act ata i Phe Thr Ile i	aaa tca ttg	tcc aga co Ser Arg Pi	ct gga cag Pro Gly Gln	ccc tgg tgt Pro Trp Cys 25	gaa gcg Glu Ala	199
cat gtc ttc His Val Phe	ttg aat aaa Leu Asn Lys 35	aat ctt t	tc ctt cag he Leu Gln	tac aac agt	gac aac Asp Asn 45	247
aac atg gtc Asn Met Val	aaa cct ctg	ggc ctc c Gly Leu L	tg ggg aag Leu Gly Lys 55	aag gta tat Lys Val Tyr	gcc acc Ala Thr 60	295
agc act tgg	qqa qaa ttg	Thr Gln T	cg ctg gga	gaa gtg ggg Glu Val Gly 75	cga gac Arg Asp	343
ctc agg atg Leu Arg Met 80	ctc ctt tgt Leu Leu Cys	Asp Ile L	aaa ccc car ys Pro Gln	Ile Lys Thr	agt gat Ser Asp	391
cct tcc act Pro Ser Thr 95	ctg caa gtc Leu Gln Val	kar atk t Xaa Xaa P 100	tt tgt caa Phe Cys Gln	cgt gaa gca Arg Glu Ala 105	gaa cgg Glu Arg	439
tgc act ggt Cys Thr Gly 110	Ala Ser Trp 115	Gln Phe A	Ala Thr Asn 120	Gly Glu Lys	Ser Leu 125	487
ctc ttt gac Leu Phe Asp	Ala Met Asn 130	Met Thr T	Trp Thr Val	Ile Asn His	Glu Ala 140	535
agt wag atc Ser Xaa Ile	Lys Glu Thr 145	Trp Lys L 1	Lys Asp Arg 150	Xaa Leu Glu 155	Xaa Tyr	583
ttc agg aag Phe Arg Lys 160	ctc tca aar Leu Ser Lys	gga gac t Gly Asp C 165	gc gat cac Cys Asp His	tgg ctc agg Trp Leu Arg 170	gaa ttc Glu Phe	631
tta ggg cac	tgg gaa gca		raa ccg ama	gtg tcm cca	rta aat	679

....

Leu Gly His Trp Glu Ala Met Pro Xaa Pro Xaa Val Ser Pro Xaa Asn	
175 180 185	
gct tca raw atc cac tgg tct tct tct art cta cca raw ara tgg atc Ala Ser Xaa Ile His Trp Ser Ser Ser Xaa Leu Pro Xaa Xaa Trp Ile	727
190 195 200 205	
atc ctg ggg gca ttc atc ctg tta vtt tta atg gga att gtt ctc atc	775
Ile Leu Gly Ala Phe Ile Leu Leu Xaa Leu Met Gly Ile Val Leu Ile 210 215 220	
tgt gtc tgg tgg caa aat ggc ara ara tcc acc tad arg tgataccacg	824
Cys Val Trp Gln Asn Gly Xaa Xaa Ser Thr Xaa Xaa 225 230	
geggegeaaa attgtteace tgtggteete gategetgae ageettgget eccaetgetg	884
tgtgttccct gagtcaagtg gaggcggagc ctgcaatgag cggaratcgc gcctctgcat	944
tocagtottg gcaacagaro aagactoogt otcaaaaaaa aaaaa	989
<210> 321	
<211> 1017	
<212> DNA	
<213> Homo sapiens	
C2137 Nome Supposed	
<220>	
<221> CDS	
<222> 3581	
<221> sig peptide	
<222> 3182	
<223> Von Heijne matrix	
score 6.69999980926514	
seq LWPFLTWINPALS/IC	
<221> polyA site	
<222> 10061016	
<400> 321	
ac atg tgc cct agt ctg gaa gag gct ccc agt gtc aag ggg act ctg	47
ac atg tgc cct agt ctg gaa gag gct ccc agt gtc aag ggg act ctg Met Cvs Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu	47
ac atg tgc cct agt ctg gaa gag gct ccc agt gtc aag ggg act ctg Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -50 -50	
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc	4 7 95
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc	
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile	95
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30	
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg qqc agg agc aga aag gtg gct cga ggt gca ccg gtc	95
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30	95
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15	95
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac	95 143
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp	95 143
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -55	95 143
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 -5 -5 -50 -50 -50 -50 -50 -50 -50 -	95 143 191
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala	95 143 191
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 -5 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala	95 143 191
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 1 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 10 15 ccc ctg ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg	95 143 191 239
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 10 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg	95 143 191 239
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 -5 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 -10 -5 -5 -6 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 -5 -5 -5 -50 -50 -50 -50 -6 -6 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7	95 143 191 239
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 1 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 10 15 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 25 30 35 gct gta ggc ccc acg gcc cca sgc	95 143 191 239 287
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 -5 -5 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 -6 -6 -5 -6 -6 -6 -5 -6 -6 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7 -7	95 143 191 239 287
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 -5 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 25 gct gta ggc ccc acg cca ggc ctc ctc cct gag gct gca gcc cca sgc Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Ala Pro Xaa	95 143 191 239 287 335
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 ccc ctg car ttg cct act gcc tgt cct ccc cac cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 25 30 35 gct gta ggc ccc acg cca ggc ctc ctc cct gag gct gca gcc cca sgc Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Pro Xaa 40 45 acg tgk ggg cac tgt tcc atc	95 143 191 239 287
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 25 30 35 gct gta ggc ccc acg cca ggc ctc ctc cct gag gct gca gcc cca sgc Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Pro Xaa 40 acg tgk ggg gca ctg ttc tca cgc agc agg cac tgg tca tgt tcc att Thr Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile	95 143 191 239 287 335
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 25 gct gta ggc ccc acg cca ggc ctc ctc cct gag gct gca gcc cca sgc Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Pro Xaa 40 acg tgk ggg gca ctg tcc tca cgc agc agg cac tgt tcc att Thr Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile 55	95 143 191 239 287 335
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 -20 -15 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 -5 1 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 10 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 25 30 35 gct gta ggc ccc acg cca ggc ctc ctc cct gag gct gca gcc cca sgc Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Pro Xaa 40 45 acg tgk ggg gca ctg tcc tca cgc agc agc agc agc gca tgt tcc att Thr Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile 55 Gtc arc tcc ccc ctc cct gtg gag acc aga arc	95 143 191 239 287 335
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu -60 -55 -50 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile -45 -40 -35 -30 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val -25 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp -10 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala 5 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg 20 25 gct gta ggc ccc acg cca ggc ctc ctc cct gag gct gca gcc cca sgc Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Pro Xaa 40 acg tgk ggg gca ctg tcc tca cgc agc agg cac tgt tcc att Thr Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile 55	95 143 191 239 287 335

70 75 80	
ttc cas aaa cat ctg ttg gtg ctg ctg gtg gct gtg gcc cat agt gtt Phe Xaa Lys His Leu Leu Val Leu Val Ala Val Ala His Ser Val 90 95	479
ctg gaa cca cct gcc ctg gtc cca aat gtg cag tgt gag atg tgc aca Leu Glu Pro Pro Ala Leu Val Pro Asn Val Gln Cys Glu Met Cys Thr	527
cac tca ggg ccc cgt gac ctg gaa gcc gca gtc gtg tcc cca gca cct His Ser Gly Pro Arg Asp Leu Glu Ala Ala Val Val Ser Pro Ala Pro	575
120 125 130 tgg gaa tgagcctgtc ctctgtgtga aggaggggt ggttctcaaa ccactgactc	631
Trp Glu ttggtgctca ggaggggcct gctgctgtcc tgggcatggg gtggtcattg ttcaagactg aggcagactc agtctttgaa agggtgcaga ggccaggcgc ggtggctcac gcctgtaatt ccagcacttt gggaggccaa ggtggacaga tcatgaggtc aggagttcga gaccagcctg gccaatacgg tgaaaccgca tctctactaa rraatawcaw aaattagtcg ggcatgggtg atgtgtgctt gtagtcccag ctactcatga ggyctgaggc agaagaatca cctgaatctg ggaggcagag gttgcagtga accaagatcg cacgactgta caccagcctg ggcgacagag tgagactccg tctcaaaaaa aaaaam	691 751 811 871 931 991
<210> 322 <211> 529	
<212> DNA <213> Homo sapiens	
<220> <221> CDS	
<222> 107427	
<221> sig_peptide <222> 107190 <223> Von Heijne matrix score 3.7999995231628 seq RFLSLSAADGSDG/SH	
<221> polyA_signal	
<221> polyA_site <222> 516529	
<400> 322	60
aaagtcagcg ctggagtcgg ctaggeggct ggaaacggcg gctgccgccg gtgactcagg gaggcgggag gccgmsggmg gagctcttcc tgcaggcgtg garacc atg gtg ctc Met Val Leu	115
acg ctc gga gaa agt tgg ccg gta ttg gtg ggg agg agg ttt ctc agt Thr Leu Gly Glu Ser Trp Pro Val Leu Val Gly Arg Arg Phe Leu Ser	163
ctg tcc gca gcc gac ggc agc gat ggc agc cac gac agc tgg gac gtg Leu Ser Ala Ala Asp Gly Ser Asp Gly Ser His Asp Ser Trp Asp Val	211
gag cgc gtc gcc gag tgg ccc tgg ctc tcc ggg acc att cga gct gtt Glu Arg Val Ala Glu Trp Pro Trp Leu Ser Gly Thr Ile Arg Ala Val	259
tcc cac acc gac gtt acc aag aag gat ctg aag gtg tgt gtg gaa ttt Ser His Thr Asp Val Thr Lys Lys Asp Leu Lys Val Cys Val Glu Phe	307
gak ggg gaa tct tgg agg aaa aga aga tgg ata gaa gtc tac agc ctt Xaa Gly Glu Ser Trp Arg Lys Arg Arg Trp Ile Glu Val Tyr Ser Leu 40 45 50 55	355

	tta	act	ctc	aat	tga	taaa	caa	gata	gtgag	ga ti	tact	catci	gg.	tacca	ac99	447
Lys	Ala	Leu 90	gcc Ala	ГÀв	Asp	Lys	Glu 95	tgg Trp	Gln	Glu	Gln	Phe 100	Leu	att Ile	Pro	392
ttt Phe	cat His	att Ile	tgg Trp 75	aaq	tat Tyr	gat Asp	aat Asn	ttt Phe 80	gct Ala	cat His	cga Arg	act Thr	gaa Glu 85	ttt Phe	Gln	344
tta	gtt Val	gga Gly	tac Tyr	tgg Trp 60	agt Ser	gta Val	kaa Xaa	ttt Phe	gga Gly 65	ggc Gly	aga Arg	atg Met	awt Xaa	aca Thr 70	gtg Val	296
ctg Leu 40	gaa	aat Asn	ttt Phe	gag Glu	aaa Lys 45	aac	gct Ala	caa Gln	ctt Leu	cgg Arg 50	aca Thr	gct Ala	cac His	tct Ser	gaa Glu 55	248
tat Tyr	gaa Glu 25	ttt Phe	cgt Arg	tct Ser	tat Tyr	tac Tyr 30	ctt Leu	aag Lys	ccc Pro	tca Ser	aag Lys 35	atg Met	aat Asn	gag Glu	ttc Phe	200
Met	tgc Cys	Ser	Ser	ttt Phe	Ala	Thr	Gly	ccc Pro	Arg	Gln	Tyr	gat Asp 20	gga Gly	ata Ile	ttc Phe	152
aga Arg	agc Ser	gcc Ala	ctg Leu	act Thr	cgg Arg	gcg Ala	ctg Leu	gcc Ala	tca Ser 1	cgg Arg	acg Thr	ctg Leu	gcg Ala 5	cct Pro	cag Gln	104
<400 aaaa	> 32 iggac	3 ac g	gctg	gctg	jc tt	ttct	cago	c gcc	gaag	ccg	cgcc	atg Met	cto Lev	gto Val	ctc Leu	56
	.> po :> 10		-													
	> po > 10	lyA_	sign	al	-											
<222		83 n He ore	ijne 5.69		8092		Ł									
<222	> CD > 45	40														
<213	> Ho	no s	apie:	ns												
<211 <212	> 32: > 10: > DN:	46 A														
	atga aaaa	cg c		ctaa	t tt	taag	tgtt	aag	catt	ttg	catt	aaaa	ta t	tcat	ataat	517 529
aag Lys	tca o Ser 1	Pro (gaa a Glu :	60 att Ile	tct Ser	tgg Trp	ggt Gly			tt t	agtt.	aaat			taat	457
		ays 1			Leu	Val	Lys	His	Asn : 65	Leu	var.	beu .	AIA	70	AI 9	

ctaggctaca	caaaactagt	tggagtgttc	cacacagagt	acggagcact	caacagagtt	627
catqttcttt	qqtqqaatqa	gagtgcagat	agtcgtgcag	ctgggagaca	taagtcccat	687
gaggatccca	gagttgtggc	agctgttcgg	gaaagtgtca	actacctagt	atctcagcag	747
aatatoctto	tgattcctac	atcgttttca	ccactgaaat	agttttctac	tgaaatacaa	807
aacatttcat	taactgctat	aggatctgtc	tgctaatggt	gcttaaattc	tcccaagagg	867
ttctcacttt	tatttgaagg	aggtggtaag	ttaatttgct	atgtttcttg	cattatgaag	927
actacateta	toctttotaa	gtaccacttc	aaaaaatakt	tctgtttact	ttctgcatgg	987
tatttcagtg	tctqtcatac	attaaaaata	cttgtcactg	tttyaaaaaa	aaaaammcc	1046
	J		-	-		

<210> 324 <211> 880 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 201..332 <221> sig_peptide <222> 201..251 <223> Von Heijne matrix score 7.80000019073486 seq VLWLISFFTFTDG/HG <221> polyA site <222> 869..880 <400> 324 60 aattgctgat ggatcagtga gcctgtgttc atgccagtga gctgctgtgg ctcagatact 120 gatactttct ttccaaacag cataagaagt gattgancca caagtatact gaaggmargg yhcccwsvar tyctggwgtg amgagataaa tcaccagtca cagactatgc acccgactgc 180 233 tgctgttcag tccagggaaa atg aaa gtt gga gtg ctg tgg ctc att tct ttc Met Lys Val Gly Val Leu Trp Leu Ile Ser Phe 281 ttc acc ttc act gac ggc cac ggt ggc ttc ctg ggg gtg agt tgg tgc Phe Thr Phe Thr Asp Gly His Gly Gly Phe Leu Gly Val Ser Trp Cys 329 Tyr Val Ser Tyr Leu Phe Ser Thr Asn Ser Pro Leu Ser Phe Arg Arg 20 15 att tagaacccct cactctctag gggactgcaa ctgcataatt taatgtactt 382 gagatcagaa gtcctgagtt ctcgtttcaa cattaccaac attcactgtg tggccttgga 442 taagtragtc atttcatctc ttcggagctt agatgatcma actgcaarag gaggatcttt 502 gattamacta tottagagat ottttocagt toaacacatg otgtactatg gottotogga 562 622 tgcagaaaaa tcacatggat ggacattagc aatccttara cactgtcttt cctgtctaca ctcgcttgag tgatgckttc atctaggatc atggttttaa tattctctac atgctgatga 682 742 ctcccagctg tatagctcca tctcagaacc tctcccctgt ccacactcac atatccatta 802 cctacgtgtt atttccagct gggaaatcca gcggaacctc ggnaacttca tttgnttcaa aatcgnaacc caatcettet tgeetatete ageaagtggt atcaetatet tteeagetae 862 880

<210> 325

<211> 1217

<212> DNA

<213> Homo sapiens

ttaggcaaaa aaaaaaaa

```
<222> 217..543
<221> sig_peptide
<222> 217..255
<223> Von Heijne matrix
      score 6.40000009536743
      seq MCLLTALVTQVIS/LR
<221> polyA_site
<222> 1206...1217
<400> 325
aatgccagtg tcagcttctc tccgaaaact gggtaatacg aaatggtctt tattggttgt
                                                                      60
gaacactcga gctgagaaac attttaggat ctttgtgtct tttgtgatga ttttgtttct
                                                                      120
                                                                      180
graagrwgga aasctgtcta aaaatattca agtgtgcaac caaggattta gatgaagcca
gcaaacaaag gaatcatgta atcaggacct gagcga atg tgc tta ctc acg gcg
                                                                      234
                                        Met Cys Leu Leu Thr Ala
                                                                      282
tta gtt aca cag gtg att tcc tta aga aaa aat gca gag aga act tgt
Leu Val Thr Gln Val Ile Ser Leu Arg Lys Asn Ala Glu Arg Thr Cys
        - 5
tta tgc aag agg aga tgg ccc tgg ngc ccc tcg ccc cgg atc tac tgc
                                                                      330
Leu Cys Lys Arg Arg Trp Pro Trp Xaa Pro Ser Pro Arg Ile Tyr Cys
                    15
                                        20
tea tee ace cea tge gat tee aaa tte eee ace gte tae tee agt gee
Ser Ser Thr Pro Cys Asp Ser Lys Phe Pro Thr Val Tyr Ser Ser Ala
                                    35
                30
cca ttc cat gcc ccc ctc ccc gtc cag aat tcc tta tgg ggg cac ccg
                                                                      426
Pro Phe His Ala Pro Leu Pro Val Gln Asn Ser Leu Trp Gly His Pro
            45
                                50
ctc cat ggt tgt tcc tgg caa tgc cac cat ccc cag gga car aat ctc
                                                                      474
Leu His Gly Cys Ser Trp Gln Cys His His Pro Gln Gly Gln Asn Leu
       60
                            65
                                                70
cag cct gcc agt ctc cad acc cat ctc tcc aag ccc aag cgc cat ttt
                                                                     522
Gln Pro Ala Ser Leu Xaa Thr His Leu Ser Lys Pro Lys Arg His Phe
ara aar aar rra tgt caa gcc tgatgaarac atgagtggca aaaacattgc
                                                                     573
Xaa Lys Lys Xaa Cys Gln Ala
                    95
aatgtacara aatgagggtt totatgotga toottacott tatcacgagg gacggatgag
                                                                     633
catasectea teccatggtg gacacecaet ggatgteece gaccacatea ttgeatatea
ccgcaccgcc atccggtcag cgagtgctta ttgtaacccc tcaatgcaag cggaaatgca
tatggaacaa tcactgtaca gacagaaatc aaggaaatat ccggatagcc atttgcctac
                                                                     813
actgggetee aaaacaeeee etgeetetee teacagakte agtgaeetga ggatgataga
                                                                     873
catgcacget cactataatg cecaeggeee cecteacace atgeageeag acegggeete
                                                                     933
teegageege caggeettta aaaaggagee aggeacettg gtgtatatag aaaageeacg
                                                                     993
gagegetgea ggattateca geettgtaga ceteggeeet eetetaatgg agaageaagt
                                                                    1053
ttttgcctac agcacggcga caatacccaa agacagagag accagagaga ggatgcaagc 1113
catggagaaa cagattgcca gtttaactgg ccttgttcag tctgcgcttt ttaaagggcc 1173
cattacaagt tatagcaaar atgcgtctag ctaaaaaaaa aaaa
                                                                    1217
```

```
<210> 326
```

<221> CDS

<211> 959

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 18..446

<221> sig_peptide

```
<222> 18..140
<223> Von Heijne matrix
      score 4.09999990463257
      seq GILILWIIRLLFS/KT
<221> polyA_signal
<222> 930..935
<221> polyA_site
<222> 948..959
<400> 326
aaaggaagcg gctaact atg gcg acc gcc acg gag cag tgg gtt ctg gtg
                                                                    50
                  Met Ala Thr Ala Thr Glu Gln Trp Val Leu Val
                      -40
                                          -35
gag atg gta cag gcg ctt tac gag gct cct gct tac cat ctt att ttg
                                                                    98
Glu Met Val Gln Ala Leu Tyr Glu Ala Pro Ala Tyr His Leu Ile Leu
-30
                   -25
                                      -20
gaa ggg att ctg atc ctc tgg ata atc aga ctt ctt ttc tct aag act
                                                                   146
Glu Gly Ile Leu Ile Leu Trp Ile Ile Arg Leu Leu Phe Ser Lys Thr
                                  -5
               -10
tac aaa tta caa gaa cga tct gat ctt aca gtc aag gaa aaa gaa gaa
                                                                   194
Tyr Lys Leu Gln Glu Arg Ser Asp Leu Thr Val Lys Glu Lys Glu Glu
                           10
ctg att gaa gag tgg caa cca gaa cct ctt gtt cct cct gtc cca aaa
                                                                   242
Leu Ile Glu Glu Trp Gln Pro Glu Pro Leu Val Pro Pro Val Pro Lys
                       25
                                         . 30
gac cat cct gct ctc aac tac aac atc gtt tca ggc cct cca agc cac
                                                                   290
Asp His Pro Ala Leu Asn Tyr Asn Ile Val Ser Gly Pro Pro Ser His
                   40
aaa act gtg gtg aat gga aaa gaa tgt ata aac ttc gcc tca ttt aat
                                                                   338
Lys Thr Val Val Asn Gly Lys Glu Cys Ile Asn Phe Ala Ser Phe Asn
               55
                                  60
ttt ctt gga ttg ttg gat aac cct agg gtt aag gca gca gct tta gca
Phe Leu Gly Leu Leu Asp Asn Pro Arg Val Lys Ala Ala Ala Leu Ala
                               75
tct cta aag aag tat ggc gtg ggg act tgt gga ccc tgt gga ttt tat
                                                                   434
Ser Leu Lys Lys Tyr Gly Val Gly Thr Cys Gly Pro Cys Gly Phe Tyr
                          90
ggc aca ttt gaa tgaaratgaa ggatcattga tttccttgtg tatggataat
Gly Thr Phe Glu
   100
ccgggaacag gccaactaaa tatttgatga atgtatgatt tcaaatacag tgaattccct
999agtcatc aaaraagacg gcattttatg gttgttttta ttaagtgtat attctttgct
cctgaaaatg ttattaaata attgtttagg ccgggcatgg tggctcatgc ctgtaatccc
agcactttca aaggetgagg caggeagate acctgaggte aggagttcaa aaccageetg
                                                                   726
gccaacatgc tgaaacctcg tctctactaa aaatacaaaa attagctggg cgtggtggtg
grtgcctgtg gtcccagctr cgtgggaggc tgaggtggga gaattgcttc aacctgggag
gcggaggttg cagtgagccg agatcatgcc actgcactcc agcctgggca acagagcaag
                                                                   906
```

<210> 327 <211> 921

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222>	29.	.724														
<221> <222> <223>	29. Von	.118 Hei re 3	1	matr 0000	9536	743 X										
<221; <222;	pol 886	yA_s	signa 91	.1												
<221: <222:	> pol	LyA_s	site 20													
<400: aagg:	> 32° agcca	7 ac g	cttt	9999	gtt	gcaa	IATE	g go et Al	g g la A	cc ac la Tì	cc ag		ga a ly T	et g hr A	at sp	52
gag Glu	ccg : Pro '	gtt Val	tcc g Ser (gly c	gag t Glu I	Leu	gtg i	-ct i	gtg Val	gca (Ala)		aca	ctt	tct Ser	ctc Leu	100
		-20	tcg ' Ser '		gly a	aac Asn	-15 art :	vet :	gac	att (gag	atq	gct	tgg	gcc	148
	- 5		atg Met	cag (Gln)	cat :	l act	aaa	gtc Val	tat Tyr	tac	aag	ctg	att	tca	tca	196
gtt Val	gac Asp	cca Pro	cag Gln	15 ttc Phe	ctg Leu	aaa Lys	ctc Leu	acc Thr	20 aaa Lys	gta Val	gat Asp	gac Asp	caa Gln 40	att	tac Tyr	244
tct Ser	gag Glu	Phe	30 cgg Arg	aaa Lys	aat Asn	ttt Phe	GIU	35 acc Thr	ctt Leu	agg Arg	ata Ile	gat Asp 55	gtg	ttg Leu	grc Xaa	292
cca Pro	gaa Glu	45 gan Xaa	ctc Leu	aag Lys	tca Ser	GIU	50 tca Ser	gcn Ala	aaa Lys	gag Glu	ccc Pro 70	cca	gga Gly	tac Tyr	aat Asn	340
tct Ser	60 ttg Leu	cca Pro	ttg Leu	aaa Lys	Leu	65 ctc Leu	gga Gly	acc Thr	ggg ggg	aag Lys 85	qct	ata Ile	aca Thr	aag Lys	ctg Leu 90	388
75 ttt Phe	ata Ile	tca Ser	gtg Val	Phe	80 agg Arg	aca Thr	aag Lys	aag Lys	gag Glu 100	aga	aag Lys	gag Glu	tca Ser	aca Thr	atg Met	436
gag Glu	gag Glu	aaa Lys	aaa Lys	95 gag Glu	ctg Leu	aca Thr	gtg Val	GIU	aaq	aag Lys	aga Arg	aca Thr	cca Pro 120	aga Arg	atg Met	484
		aga Arg	110 aag Lys	~~~	ata	a+a	gtg Val	TID	aaq	aaa	aag	agg	aag Lys	gaa	tca Ser	532
aca Thr	Glu	Lys		aaa Lys	ctg Leu	aca Thr	гÀг	gag Glu	gag Glu	aaa Lys	aag Lys 150	gga Gly	aag	aag Lys	ctg Leu	580
Thr	Lys		tca Ser	aca Thr	Lys	gtg Val	ata	aaa Lys	aag Lys	cta Leu 165	tgt Cys	aag	gta Val	tac Tyr	agg Arg 170	628
155 gaa Gly		cac His	tct Ser	Arg	Ser	+ = +	gac Asp	tca Ser	att : Ile 180	gag Glu	act	aca Thr	agt Ser	aco Thi	acg Thr	676
gte Val	g cta L Lev	cti Lei	ı Ala	Gln		cct	tto Lev	g gtt 1 Val	aaa Lys	a tat	aaa Lys	tto Phe	tts Lev 200	2	c aat r Asn	724
tg:	aagga ttgt:	atac atga	190 gcag actg		ac a aa a	tctt igact	tcta	ar to	taa	cagto	agg agtt	gagct cgatt	tgct	ctg	gtcattc ggtatac	784 844

904

tgtttcttgg ctgacactac tggtcaagta agaaatttgt aaataaattt cttttggttc

921 ttattaamaa aaaaaas <210> 328 <211> 1344 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 404..586 <221> sig_peptide <222> 404..466 <223> Von Heijne matrix score 4.09999990463257 seq SLMFFSMMATCTS/NV <221> polyA_signal <222> 1304..1309 <221> polyA_site <222> 1334..1344 <400> 328 ataatttaat gcaaaatatc cttttatgaa tttcatgtta atattgtgaa atattaaaat aattccacaa tagttgagaa aaatgagcat ttttttccat ttttaaaaaa tgcatagaaa 120 agacaatttt aaaatcctgg gamccawatt tatttagaag tagctgttag taaaacatta 180 gaaaaggagt caggccatba ggttatttat nbnaatctct aagcaattag gntgaagtta 240 ttaagtcaag cctagaaaag ctgcctcctt gtaaggcttt catgacaatg tatagtaatc 300 breagtgtee aattettege acteeteagg aatateacta ceteaggtta eggtacaeag 360 gctataattg atgatgatgt tcagataact gaagacacaa taa atg aca ttc aga 415 Met Thr Phe Arg cat cag gac aat too oto atg tto ttt tot atg atg goo acc tgt acc 463 His Gln Asp Asn Ser Leu Met Phe Phe Ser Met Met Ala Thr Cys Thr -5 -10 -15 age age gtg ggt tte ace cae aca acg atg age tgt tet ett act tet 511 Ser Asn Val Gly Phe Thr His Thr Thr Met Asn Cys Ser Leu Thr Ser cca gtt gat ttt aaa gac ttg tta aga gtc tta cta ata aaa ttt ggg 559 Pro Val Asp Phe Lys Asp Leu Leu Arg Val Leu Leu Ile Lys Phe Gly 25 020 606 tat gat aga aaa too aca ato aaa tot tgaaccaaat aacatattaa Tyr Asp Arg Lys Ser Thr Ile Lys Ser 35 666 attactaata tttaagtgat ggaagacaca caaaaaactt aaaagcacga acaacctaac ttgaaaaara attttaaaat atgattaacc tgaaraaaar araatcctaa ragccaaagc tectttttat ttagettgga atttteetat tggtteetaa caaactgtee caatgteata taaggaaaca tgatctatta cattccttta taacaacgtg gararactat aaacctatgt 846 aagtagtaaa actatatcag adactcagga ractgactww aaggcctgga tctgcagtgt attatctgta taaaaattgg cagggggaag ctaaaaggaa aggagattgg agatctcaat 966 totatcatgg tgtatttcat acgcaaatca ragcatgcat tgttttttgt ttttggaaar 1026 avaarggaag tgtgttctgc cccatgtttc cttccgtgtt tatagttcaa actctatata tacttcaggt attttttgtt tagcccttca ttataaatgg gcaggaaatt gtttatcaac ctagccagtt tattactagt gaccttgact tcagtatctt gagcattctt ttatattttt 1206 cttttattat cctgagtctg taactaaaca attttgtctt caaattttta tccaatatcc 1266 attgcaccac accaaatcaa gcttcttgat tttcaaaaat aaaaaggggg aaatacttac 1326 1344 aacttgtaaa aaaaaaaa

<210> 329 <211> 585 <212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 331..432

<221> sig_peptide

<222> 331..387

<223> Von Heijne matrix
 score 7
 seq AGLSSCLLPLCWL/ER

<221> polyA signal

<222> 548..553

<221> polyA_site

<222> 573..585

<400> 329
aagcctaggt gtggcgccc gaccggactt tcacttctgg ccagcccttt ccccacctgg
gcgcgggass ggtgccagtc tttaaacaac ctctcgatgg gtcccacgaa gatgtttcca 120
gacccttgga atgccaagtt caagtttagc tatgtctcgc ggagaggccg gtggaagaag 180
caacgagaat gaagcaccc agttctctgc tgagcacatg ggcatctgca ataaagattt 240
aatttcccag cttctcctga agctcggtat ggccacaaca ctaaattctg cccgaggaga 300
attgagcaaaa tagtatggga cttccaagaa atg ttt tta aag tca ggg gca ggc
Met Phe Leu Lys Ser Gly Ala Gly
-15

ctt tct tca tgc ctt ctt cct ctt tgc tgg ctg gaa cgc aaa gac cat

402
Leu Ser Ser Cys Leu Leu Pro Leu Cys Trp Leu Glu Arg Lys Asp His

-10 -5 1 5
ggc agg agg cca agc asc cat cct gga agg tgaaagcctc atactaagga 452

Gly Arg Arg Pro Ser Xaa His Pro Gly Arg

cgtcaracag cgaaataara rcctgggtcc ttgaccctgt aaasatctcc ctccccatcc 512 tggtctgtct gccttgactc ctttcatatg aaaaaaaataa acttttaact tgcgtwaacc 585

<210> 330

<211> 914

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 59..703

<221> sig_peptide

<222> 59..220

<223> Von Heijne matrix
score 5.09999990463257
seq FLLSQMSQHQVHA/VQ

<221> polyA_site <222> 903..914

<pre><400> 330 acaaatatca atgatgttta tgaatctagt gtgaaagtkt taatcacatc acaaggct 58</pre>																
acaa	atat	ca a	itgat	gttt	a tg	gaato	tagt	gto	aaag	jtkt	taat	caca	tc a	t+a	ger Jgcr	106
atg	aac	rra	tat	gca	agt	cca	ttc	aac	tgw	Caa	ttg	Yaa	Tyr	Len	Yaa	100
Met	Asn	Xaa	Tyr		Ser	Pro	Pne	ASI	-45	GIII	ьеи	Aaa	TYL	-40	21.00	
			ttc	-50		~+~	cat	202		aas.	aga	ota	att		ctq	154
ttg	agc	agr	Phe	gag	Cyc	Mal Gri	Hie	Ara	Asp	Glv	Ara	Val	Ile	Thr	Leu	
ren	ser	Arg	-35	GIU	Cys	Val	1115	-30		-	5		-25			
t a t	+ = +	cad	gag	cad	gag	cta	caq		ttt	ctt	ctq	tct	cag	atg	tca	202
COT	Tar	Gln	Glu	Gln	Glu	Leu	Gln	Asp	Phe	Leu	Leu	Ser	Gln	Met	Ser	
		-20					-15					-10				
cag	cac	cad	gta	cat	qca	gtt	cag	caa	ctc	gcc	aag	gtt	atg	ggc	tgg	250
Gln	His	Gln	Val	His	Ala	Val	Gln	Gln	Leu	Ala	Lys	Val	Met	Gly	пр	
	- 5					1				5					10	
caa	gta	ctg	agc	ttc	agt	aat	cat	gtg	gga	ctt	gga	cct	ata	gag	agc	298
Gln	Val	Leu	Ser	Phe	Ser	Asn	His	Val	Gly	Leu	Gly	Pro	Ile	Glu	Ser	
				15					20					25	-t-a	346
abt	ggt	aat	gca	tct	gcc	atc	acg	gtg	gcc	CCC	caa	gtg	gtg	Thr	Mot	340
Xaa	Gly	Asn	Ala	Ser	Ala	Ile	Thr	Val	Ala	Pro	GIn	vai	40	TIII	Mec	
			30					35	- 4			202		taa	++-	394
cta	ttt	cag	ttc	gta	atg	gac	ctg	aaa	grg	gça	90a	Ara	Len	Trn	Phe	
Leu	Phe		Phe	Val	Met	Asp		гур	Val	Hia	Ala	55	шси			
		45	gta			~+ ~	50	3.00	++-	caa	222		ato	ttt	tac	442
agt	ttc	CTC	gta Val	acc	Aat Aen	y La	Tave	Thr	Phe	GÌn	Lvs	Val	Met	Phe	Tyr	
Ser		Leu	vai	1111	ASII	65	пyз	1111	1110		70				•	
222	60 ata	202	aat	ada	atc	atc	ttc	ata	qqq	cat	tca	aar	aag	ttc	agt	490
Lve	Tle	Thr	Asn	Glv	Val	Ile	Phe	Val	Gly	His	Ser	Lys	Lys	Phe	Ser	
75					80					85					90	
aas	ata	aaa	taa	aag	gtc	kaa	att	ttg	ttt	ata	aaa	tgg	arm	tgc	tta	538
Gly	Ile	Lys	Trp	Lys	Val	Xaa	Ile	Leu	Phe	Ile	Lys	Trp	Xaa	Cys	пеп	
				95					100					102		F 0.6
tgt	ctg	cac	tta	gcc	ctt	gtc	tac	tat	gat	ttt	tto	car	atg	ttt	cct	586
Cys	Leu	His			Leu	Val	Tyr	Tyr	Asp	Phe	Phe	GIn	Met	Pne	Pro	
			110					115				. ~~~	120		tat	634
aaa	raa	gtt	tcc	ara	aac	ttt	gac	ttg	aaa	tgt	ttg	Car	Tle	Acn	tat Tvr	051
Lys	Xaa			Xaa	Asn	Pne	Asp	ьeu	. гу	cys	ье	135	. 110		Tyr	
		125				20+	130			ato	cto			aaa	ata	682
aag	cac	aaa	gaa	gar	ata Tla	. act	SAY	Twe	ayo	val	Lei	Phe	Leu	Lys	Ile	
гÀг	140		G L U	GIL	1 116	145		. Dys		, , , ,	150	,		•		
	140		j aaa	+~+				rcact	ttc	aaac			ttta	taaa	it	733
Tle	Tla	Aye	, aaa , Lys	Cve	. Dhe	Tle		, •								
155			, Lys	. Cys	160											
gac	aagt	act	ttaa	aato	ica c	raaqt	ttat	g ta	cagt	tgta	a tat	acag	ıtat	gaca	agatgt	793
222	ataa	tat	attt	ttca	ita c	agtt	taaa	aa ta	ittad	ctaac	: tta	aggg	ittt	ctat	gtgctt	853
ttt	aaaa	tat	tect	tctt	tg a	tgtt	gaca	at ca	aata	aaagt	ato	tggt	tta	aaaa	aaaaaa	913
a					_	-	-									914

<210> 331

<211> 1161

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 672..752

<221> sig_peptide <222> 672..722 <223> Von Heijne matrix score 4.30000019073486 seq LLYAHLSFTSKRA/VV <221> polyA_site <222> 1150..1161 <400> 331 aagatatcac tgtcttgttt tcacttagat cctacttaca aagtgagggt tattaacaga ataaagcctt cctttaaagc tttataataa tcatatttat taataatgct gttgtgcata 120 cttatagtat gcatatattc agcatatgtt gcatgtsttc agaattacat aagatgaaat 180 240 ccctttcatt gcaacttgca agtgagaaaa gatccttagt ggctctggtg gaagaaatag tatttettet tetcagggtg tetceetgee ttggeceete ecagaageee eggetttaaa 300 agtgaaaatg tttgaaacat gaaacatgtc tgtaggaagc atcagcatgg ccataagtgc 360 artgattttc atatatgcct ctgcccattt caaatatatt tttgacatga ataaatctaa 420 cagtatacar aataattcat gtaaraccct aacgtgtaca tgtgaaaaag catttctata 480 taatgtgagg agcactggcc atcaattagg gaaataaagg tcatgtaata ttgcaaattt 540 tcaaaataga gcsstgcaag ataactgcaa tcataccaaa aactatttga gtaaatggat 600 ttttaaagta atttttgttt aaaaaaattt atatttcaga agsagaaaat gtcaaatgat 660 agtetttgta a atg gtg gtg cac ett ete tat gea eat etg tet ttt aca 710 Met Val Val His Leu Leu Tyr Ala His Leu Ser Phe Thr -10 -15 752 tca aaa aga gct gtg gtc atg cta aaa tta gag ata act ttt Ser Lys Arg Ala Val Val Met Leu Lys Leu Glu Ile Thr Phe 5 1 tgaatgactt ggtcaagctg tgtgtaaaat atttaaccat aagtcaagta cagtgtacta 812 872 tgtttaataa agttacattt aatgcattta ttgcatatat gaatatatac atgaagaggc tttatgtctt ctggtatttg attttgaatg ttttttaagt cagtggtgcc tttaggcaag 932 aactttcgaa attaatcatt ctttgtgttt tctgattttt caggtaacat gtacactatt tagaaaccat catagtttat tcaccttaaa aaattgattg tattatttaa atatatcact 1052 tagatgggca tttcctataa ttaggatatt ccaaatagtt gctgaaatca attgtgccat 1112 1161 tgaccaatgg atgcacttgg ttagccttaa ttttttyaaa aaaaaaaaa

<210> 332 <211> 363 <212> DNA

<213> Homo sapiens

<220>
<221> CDS
<222> 57..311

<221> sig_peptide

<222> 57..128

<223> Von Heijne matrix
score 5.30000019073486
seq LFHLLFLPHYIET/FK

<221> polyA_signal <222> 332..337

<221> polyA_site <222> 351..363

<400> 332
acatttetta etgeettaeg eteateetga ggteeacett ggtetetaaa aacace atg

	107
tgt tct cat gcc tcc atg tct ttt cac aca ctg ttc cat ttg ctc ttc Cys Ser His Ala Ser Met Ser Phe His Thr Leu Phe His Leu Leu Phe -20 -15 -10	107
ctc cca cat tac att gaa act ttc aag cct cag tcg aaa cat tgc ttc Leu Pro His Tyr Ile Glu Thr Phe Lys Pro Gln Ser Lys His Cys Phe	155
ttc tgg ata gca gcc ttc ttg aca tcc ctc ctc act ccc cag tcc cta Phe Trp Ile Ala Ala Phe Leu Thr Ser Leu Leu Thr Pro Gln Ser Leu	203
10 15 20 25 cag ggc ttc cat agc tct tta tgt gca ctt cga tcc cag cat ttt cca Gln Gly Phe His Ser Ser Leu Cys Ala Leu Arg Ser Gln His Phe Pro	251
tcg act tgt aat tgt ttc tgc tac ctg aca atc atc gcc ttg drd tac Ser Thr Cys Asn Cys Phe Cys Tyr Leu Thr Ile Ile Ala Leu Xaa Tyr	299
45 50 55 tgg gac aac ctt tgattactca ttatatcctc aataaatatt tgttgaacca	351
Trp Asp Asn Leu 60	363
aaaaaaaaa aa	303
<210> 333	
<211> 645 <212> DNA	
<213> Homo sapiens	
<220> <221> CDS	
<222> 80232	
<221> sig_peptide	
<222> 80127 <223> Von Heijne matrix	
score 3.7000004768372 seq IALTLIPSMLSRA/AG	
<221> polyA_signal	
<221> polyA_site <222> 634645	
<400> 333	
accttcttgt tatttatgct attctctttg tggctccatt cttcttcaa tcttctcagc ttataaccgt ctttccctt atg cta agg ata gcc ctt aca ctc atc cca tct Met Leu Arg Ile Ala Leu Thr Leu Ile Pro Ser -15	60 3112
atg ctg tca agg gct gct ggt tgg tgc tgg tac aag gag ccc act cag	160
Met Leu Ser Arg Ala Ala Gly Trp Cys Trp Tyr Lys Glu Pro Thr Gln -5 1 10	
cag ttt tct tac ctt tgc ctg ccc tgc ctt tca tgg aat aar aaa ggc Gln Phe Ser Tyr Leu Cys Leu Pro Cys Leu Ser Trp Asn Lys Lys Gly	208
aac gtt ttg cag ctt cca aat ttc tgaaraaact aatctcarat tggcagttaa Asn Val Leu Gln Leu Pro Asn Phe	262
30 35 agtcaaaatg ttgccaaata tttattcctt ttgcctaakt ttggctaccc ggttcaattg	322
ctttttattt ttaatgtett gaetettear agttegtace teaaaaraae aatgaraaea tttgetttge tttetgetga atecetaate teaacaatet atacetggae tgteeagtte	382 442
tectectgtg ctatettete ttetatecaa gtaraatgta ygecaggare teetteeete tareaattte taetaaaatg teeaagtara atgttteett ttacaateaa attactgtat	502 562

ttattaattt gctaraatcc aktaaatcat tttggtagct ctggctgtgc tatcaataaa aagatgaaag caaaaaaaaa aaa	622 645
<210> 334 <211> 400 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 91291	
<pre><221> sig_peptide <222> 91219 <223> Von Heijne matrix</pre>	
<221> polyA_signal	
<221> polyA_site <222> 389400	
<400> 334 aacaaaagga gagttttata attcacttta aaaggagatt tgatggtaaa gtttaaagat taaaatattt tgttcttcaa ttacagagcg atg acc cca cag tat ctg cct cac Met Thr Pro Gln Tyr Leu Pro His -40	60 114
ggt gga aaa tac caa gtt ctt gga gat tac tct ttg gca gtg gtc ttc Gly Gly Lys Tyr Gln Val Leu Gly Asp Tyr Ser Leu Ala Val Val Phe -35 -20	162
ccc ctg cac ttt tct gat cta att tct gtt tta tac ctt ata ccc aaa Pro Leu His Phe Ser Asp Leu Ile Ser Val Leu Tyr Leu Ile Pro Lys -15 -10 -5	210
aca ctt act acc aac aca gct gtt aaa cat tct ata caa aaa aat tgt Thr Leu Thr Thr Asn Thr Ala Val Lys His Ser Ile Gln Lys Asn Cys 1 5 10	258
atg mat ctg gta tta gga aaa tta ctt tca cag taaatatcaa agaaaaaaga Met Xaa Leu Val Leu Gly Lys Leu Leu Ser Gln	311
ttaagggtet etttgecatg etttteatea tatgeaceaa atgtaaattt tgtaeaataa aattttattt eetaagyaaa aaaaaaaaa	371 400
<210> 335 <211> 496	
<212> DNA <213> Homo sapiens	
<220>	
<221> CDS <222> 196384	
<221> sig_peptide <222> 196240 <223> Von Heijne matrix score 6.69999980926514	

seq ILSTVTALTFARA/LD

<221> polyA_signal	
<221> polyA_site <222> 485496	
<pre><400> 335 aaaaaattgg tcccagtttt caccctgccg cagggctggc tggggagggc agcggtttag attagccgtg gcctaggccg tttaacgggg tgacacgagc htgcagggcc gagtccaagg cccggagata ggaccaaccg tcaggaatgc gaggaatgtt tttcttcgga ctctatcgag gcacacagac agacc atg ggg att ctg tct aca gtg aca gcc tta aca ttt</pre>	60 120 180 231
gcc aga gcc ctg gac ggc tgc aga aat ggc att gcc cac cct gca agt Ala Arg Ala Leu Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser 1 5 10	279
gag aag cac aga ctc gag aaa tgt agg gaa ctc gag agc agc cac tcg Glu Lys His Arg Leu Glu Lys Cys Arg Glu Leu Glu Ser Ser His Ser	327
gcc cca gga tca acc cag cac cga aga aaa aca acc aga aga	375
30 35 40 45 tot toa goo tgaaatgaak cogggatcaa atggttgctg atcaragccc	424
Ser Ser Ala atatttaaat tggaaaagtc aaattgasca ttattaaata aagcttgttt aatatgtctc	484
aaacaaaaaa aa	496
<pre><210> 336 <211> 968 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 54590 <221> sig_peptide <222> 54227 <223> Von Heijne matrix</pre>	E.G.
atatttgccc cttactttat cttgtgcctt gagaaattgc tggggagaga ggt atg Met	56
tcc act ggg cag ctg tac agg atg gag gat ata ggg cgt ttc cac tcc Ser Thr Gly Gln Leu Tyr Arg Met Glu Asp Ile Gly Arg Phe His Ser -55 -50 -45	104
cag cag cca ggt tcc ctc acc cca agc tca ccc act gtt ggg gag att Gln Gln Pro Gly Ser Leu Thr Pro Ser Ser Pro Thr Val Gly Glu Ile -40 -35 -30	152
atc tac aat aac acc aga aac aca ttg ggg tgg att ggg ggt atc ctt Ile Tyr Asn Asn Thr Arg Asn Thr Leu Gly Trp Ile Gly Gly Ile Leu -25 -10	200
atg ggt tot ttt cag gga acc att gct gga caa ggc aca gga gcc acc Met Gly Ser Phe Gln Gly Thr Ile Ala Gly Gln Gly Thr Gly Ala Thr	248

0

WO 99/31236

				-5					1				5				
tcc	att	tct	gag	ctc	tgc	aag	gga	caa	gaa	cta	gag	cca	tca	999	gct	. 2	296
Ser	Ile	Ser	Glu	Leu	Cys	Lys	Gly	Gln	Glu	Leu	Glu	Pro	Ser	Gly	Ala		
		10					15					20				-	344
aāa	ctc	act	gtg	acc	cca	CCC	caa	gcc	gtc	agc	Leu	Gln	Glv	atc Ile	Tvr	_	
GIA		Thr	vaı	Ala	PIO	30	GIII	ATG	Val	361	35	O+	,	Ile	•		
300	25 Ct G	cct	taa	cta	cta		ctt	ttt	cac	tcc		gcc	cta	rgg	gna	3	392
Thr	Leu	Pro	Trp	Leu	Leu	Gln	Leu	Phe	His	Ser	Thr	Ala	Leu	Xaa	Add		
40					45					50					55		
dtt	cag	caa	cct	aat	gga	tct	cta	tct	ctg	aac	atc	tct	tca	tcc	cat	4	440
Xaa	Gln	Gln	Pro		Gly	Ser	Leu	Ser	Leu	Asn	Ile	Ser	Ser	Ser 70	HIS		
				60				ata	65	cc =	~~=	atá	gac		acc	4	488
gct	ccr	rgt	cca	rca	acc Th~	Cyre	Thr	Len	Glu	Pro	Glv	Val	Asp	cct Pro	Thr		
Ala	Pro	Ada	75	Aaa	1111	Cys		80			- -1		85				
cga	sct	atc	tat	att	aat	ccc	cat	ccc	cca	cca	cca	atc	tta	aaa	abc	1	536
Arg	Xaa	Val	Cys	Ile	Asn	Pro	His	Pro	Pro	Pro	Pro	IIe	Leu	Lys	Xaa		
_		90					95					100				1	584
cct	ctg	tcc	CCC	tac	cct	aaa	CCC	cag	tta	ggt	acc	Cat	gct	999 G) v	Gln	•	204
Pro		Ser	Pro	Tyr	Pro	Lуs 110	Pro	GIN	ьец	GIY	1115	nis	AIG	Gly	0211		
	105	+ > >	caat:	++=	tacai		ta c	tagt	ttta	t ta		accq	ttc	cagg	gta		640
	Asn		caac	cca	cgcu.	-~55	. .	5-				_			_		
120																	
gct	ttga	aaa	aagt	atct	ca a	aaag	gcaa	c at	gggc	cgag	cgc	agtg	gct	cacg	cctgta		700 760
atc	ccaq	cac	tttg	qqaq	gc c	aagg	tggg	c ag	atcg	cctg	agg	tctg	gag	ttca	agacc	a	820
gcc	tggc	caa	cagg	gtga	aa c	cccg	tctc	t ac	aaaa	atar	gaa	aatt	rgc	catg	tgtgg1	-	880
ggc	agac	gtc	tgtr	gtcc	ca g	ctat	ccag	g ag	ccac	taca	ctc	cago	cta	aaca	aacccaga	-	940
gga	tgcg	gag	gttg	cagt	ga g aa a	aaaa	mcm	9 -9	CCac	-9-9		50	,	JJ J	•	-	968
Lyy	Latt	ccg		uaaa	~ CL CL												

<210> 337 <211> 901

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 133..846

<221> sig_peptide

<222> 133..345

<223> Von Heijne matrix
 score 9.39999961853027
 seq VVSFLLLLAGLIA/TY

<221> polyA_site <222> 890..901

<400> 337 aagcagette caggateetg agateeggag cageegggt eggagegget ceteaagagt 60 tactgatcta tnnatggcag agaaaaaaa attgtgacca gagacgtgta gcaatgaaca 120 aggaacrtca ta atg rwn nnk ttc aca gac ccc tct tca gtg aat gaa aag 171 Met Xaa Xaa Phe Thr Asp Pro Ser Ser Val Asn Glu Lys -65 -70 aag agg agg gag cgg gaa gaa agg cag aat att gtc ctg tgg aga cag 219 Lys Arg Arg Glu Arg Glu Glu Arg Gln Asn Ile Val Leu Trp Arg Gln -50 -55 ccg ctc att acc ttg cag tat ttt tct ctg gaa atc ctt gta atc ttg 267 WO 99/31236

```
Pro Leu Ile Thr Leu Gln Tyr Phe Ser Leu Glu Ile Leu Val Ile Leu
                            -35
aag gaa tgg acc tca aaa tta tgg cat cgt caa agc att gtg gtg tct
                                                                      315
Lys Glu Trp Thr Ser Lys Leu Trp His Arg Gln Ser Ile Val Val Ser
                                            -15
                        -20
ttt tta ctg ctg ctt gct ggg ctt ata gct acg tat tat gtt gaa gga
                                                                      363
Phe Leu Leu Leu Ala Gly Leu Ile Ala Thr Tyr Tyr Val Glu Gly
                                        1 .
                    -5
                                                                      411
gtg cat caa cag tat gtg caa cgt ata gag aaa cag ttt ctt ttg tat
Val His Gln Gln Tyr Val Gln Arg Ile Glu Lys Gln Phe Leu Leu Tyr
            10
gcc tac tgg ata ggc tta gga att ttg tct tct gtt ggg ctt gga aca
                                                                      459
Ala Tyr Trp Ile Gly Leu Gly Ile Leu Ser Ser Val Gly Leu Gly Thr
                            30
ggg ctg cac acc ttt ctg ctt tat ctg ggt cca cat ata gcc tca gtt
                                                                      507
Gly Leu His Thr Phe Leu Leu Tyr Leu Gly Pro His Ile Ala Ser Val
                        45
    40
aca tta gct gct tat gaa tgc aat tca gtt aat ttt ccc gaa cca ccc
                                                                      555
Thr Leu Ala Ala Tyr Glu Cys Asn Ser Val Asn Phe Pro Glu Pro Pro
                    60
tat cot gat cag att att tgt cca gat gaa gag ggc act gaa gga acc
                                                                      603
Tyr Pro Asp Gln Ile Ile Cys Pro Asp Glu Glu Gly Thr Glu Gly Thr
                                    80
                75
att tot ttg tgg agt atc atc tca aaa gtt agg att gaa gcc tgc atg
                                                                      651
Ile Ser Leu Trp Ser Ile Ile Ser Lys Val Arg Ile Glu Ala Cys Met
                                                     100
                                95
            90
tgg ggt atc ggt aca gca atc gga gag ctg cct cca tat ttc atg gcc
                                                                      699
Trp Gly Ile Gly Thr Ala Ile Gly Glu Leu Pro Pro Tyr Phe Met Ala
                                                115
                            110
aga gca gct cgc ctc tca ggt gct gaa cca gat gat gaa gag tat cag
                                                                      747
Arg Ala Ala Arg Leu Ser Gly Ala Glu Pro Asp Asp Glu Glu Tyr Gln
                                            130
                        125
                                                                      795
gaa ttt gaa gag atg ctg gaa cat gca gag tct gca caa gta aga aca
Glu Phe Glu Glu Met Leu Glu His Ala Glu Ser Ala Gln Val Arg Thr
                    140
                                         145
gtg ggg ata gaa aat aga aca ctt tac ttc cta aag agg cta tta
                                                                      843
Val Gly Ile Glu Asn Arg Thr Leu Tyr Phe Phe Leu Lys Arg Leu Leu
                                     160
                155
 agg taaaattgtt agtagttact ctgaagaaga aaactgctaa agtaaaaaaa aaaaa
                                                                      901
 Arg
```

<210> 338

<211> 1347

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 138..671

<221> sig_peptide

<222> 138..248

<223> Von Heijne matrix score 3.5 seq LVFNFLLILTILT/IW

<221> polyA_signal

<222> 1319..1324

<221> polyA_site

<222> 1338..1347

To the transfer of

<400> 338							•	
aagaatgctt	atasatsa	c aactaaa	ata aca	atatttc	ttctqaaa	tt ctca	ggcagt	60
cagactgtct	tagggagtag	c ttgataa	aat ago	ccttatc	caggittt	ta tcta	aggaat	120
cccaagaaga	caggcaaac	ta ded ec	ra cao t	ca agg c	tt atg t	ca qaa	aag	170
cccaagaaga	ctgggga a	icy gag ag let Glu Ar	ga cag c	er Arg I	7al Met S	er Glu	Lys	
	M			er Arg (-30			
			35	~_~		ctt ata	ttc	218
gat gag tat	cag ttt	caa cat c	cag gga	aca aca	gag Ctg	Lou Val	Phe	
Asp Glu Tyr	Gln Phe	Gln His C	GIN GIA	Ala Val	GIU Leu	Leu val	. Phe	
-25		-20			-15			266
aat ttt ttg	ctc atc	ctt acc a	att ttg	aca atc	tgg tta	ttt aaa	aat	266
Asn Phe Let	Leu Ile	Leu Thr 1	Ile Leu	Thr Ile	Trp Leu	Phe Lys	s Asn	
-10		- 5		1		5		
cat cga tto	cqc ttc	ttg cat g	gaa act	gga gga	gca atg	gtg tat	: aāc	314
His Arg Phe	Ara Phe	Leu His	Glu Thr	Gly Gly	Ala Met	Val Ty	Gly	
	N 10		15			20		
ctt aya atq	r dda cta	att tta o	csa tat	gct aca	gca cca	act gat	att	362
Leu Xaa Met	Cly Leu	Tle Leu 3	Xaa Tvr	Ala Thr	Ala Pro	Thr Asp	o Ile	
Deu Maa Met	. Gly neu	110 200 -	30		35			
gaa agt gg:				aaa cta	act ttc	agt cca	a tca	410
Glu Ser Gly	rot gud	Tue Ace (Cyc Val	Lve Len	Thr Phe	Ser Pro	Ser	
	xaa vai		cys vai	пуз пец	50			
40		45		~++ +-+		saa ta	aar	458
act ctg ctg	g gtt aat	atc act	gac caa	get tat	Glu Tur	LVE TV	r Ivs	• • •
Thr Leu Lei	ı Val Asn		Asp Gin	var fyr	GIU IYI	пур ту.	70	
55		60		65				506
aga gaa ata	a agt cag	cac amc	atc aat	cct cat	cam gga	aat gt	z ala	500
Arg Glu Ile	e Ser Gln	His Xaa	Ile Asn	Pro His	xaa Giy	ASH AI	a 116	
	75			80		85		E E 1
ctt gaa aa	g atg aca	ttt gat	cca raa	atc ttc	ttc aat	gtt tt	a ctg	554
Leu Glu Ly	s Met Thr	Phe Asp	Pro Xaa	Ile Phe	Phe Asn	vai ne	n ren	
	90		95			100		
cca cca at	t ata ttt	cat gca	gga tat	agt cta	aag aag	aga ca	c ttt	602
Pro Pro Il	e Ile Phe	His Ala	Gly Tyr	Ser Leu	Lys Lys	Arg Hi	s Phe	
10			110		115			
ttt caa aa	e tta gga	tct att	tta acg	tat gcc	ttc ttg	gga ac	t gcc	650
Phe Gln As	n Leu Gly	Ser Ile	Leu Thr	Tyr Ala	Phe Leu	Gly Th	r Ala	
120	200 027	125		•	130			
atc tcc tg	c atc atc		taaqtqa	cat tcgg	agetea a	gttgcag	gt	701
Ile Ser Cy	c Tle Val	Tle Gly			J	-		
	S IIC VAI	140						
135 ggctgtgggg	++ + -	et etetes	aaaa ta	taacactt	ccaggat	tct tqc	taackaa	761
gaaaattgtc	teygtgate	ce gegega	aggga cc	ttatatat	tttttcw	gac tta	attccac	821
gaaaattgtc		ar tawacc	ataw at	cegeacge	waggatt	act aga	atttata	881
ggcttckgam	aaatacaa	gg cttcaa	latea aa	gcaaacta	waggatt	act tat	atassac	941
tgtgagttct	ggacttct	ga cttagg	gaat gt	ggattatt	tgccttg	age cae	ttttata	1001
gcattgcatt	cttctttt	ag tttgag	taat sc	cgatatgo	ccactgc	acc 566	ccaaca	1061
fratattgag	agacetta	cc tqtatt	tggc ag	gagtgcaa	aagtaac	tat aty	Ccaagag	1121
ttttcttct	aaaqqaaa	gt ttacaa	igaca go	agtctgaa	acagata	tgt cca	aatatta	
acagagttgg	ttaataca	gg gatago	tttt ca	.gttaatac	cctgtag	aat gca	gaetett	1181
tttttcatto	tattttct	to attato	ictac tg	agccctaa	. gtcacac	gtt ata	tactetg	1241
gcttgcagct	catcataa	ag taaaat	gtgg ta:	ccaaatgg	tgaaggc	aat cca	gcctctg	1301
ataatcccgt	ccaataca	tt aaagct	ccac tg	caggaaaa	aaaaaa			1347
		_						

<210> 339

<211> 987

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

PCT/IB98/02122

<222> 124..411 <221> sig_peptide <222> 124..186 <223> Von Heijne matrix score 6.30000019073486 seg MVALCCCLWKISG/CE <221> polyA_signal <222> 948..953 <221> polyA_site <222> 971..983 <400> 339 aagacgctgc ctttagggag agataaaaag cataatgaca ttagctagga aagttaattt tcagttctta ctgaagtgct gtatgaaact gaaatttcca aggaactgaa ttttgtgagc 120 168 caa atg agc atg caa ttc ttg ttt aag atg gtg gcc tta tgc tgt tgt Met Ser Met Gln Phe Leu Phe Lys Met Val Ala Leu Cys Cys -15 ctc tgg aag atc tcc ggc tgt gag gaa gtc cct cta act tac aac ctg 216 Leu Trp Lys Ile Ser Gly Cys Glu Glu Val Pro Leu Thr Tyr Asn Leu - 5 264 ctc aag tgc ctc cta gat aaa gcg cac tgt gta ctc ctg aca cct tgt Leu Lys Cys Leu Leu Asp Lys Ala His Cys Val Leu Leu Thr Pro Cys 20 15 312 ggt tac atc ttt tcc ttg atc agt cca gaa att ctc aaa ctc act tta Gly Tyr Ile Phe Ser Leu Ile Ser Pro Glu Ile Leu Lys Leu Thr Leu 35 30 atc act ttg.cav atc ctc tta ata ctc aaa aat cta cac tta ctg tgg 360 Ile Thr Leu Xaa Ile Leu Leu Ile Leu Lys Asn Leu His Leu Leu Trp 50 ctg aca gtt tca agc awa tgt gtt cat cgc agt agt gca aga aaa gaa 408 Leu Thr Val Ser Ser Xaa Cys Val His Arg Ser Ser Ala Arg Lys Glu 65 461 aag tagaagaacc ctgcagagat ttgatggaac ccagcttcta ttcattaaaa Lys 75 ccaatggcaa aatataaagc aaataggagg tgacgaaggt tacaaaaata cgtattgttt 521 atgttttccc tggggtgtgc tgattgtcag gcatcagttc cctgtgccat tcattcccca 581 641 acacagcatg catcagaaat tttatcaata aatgctttct ctctcaatgt tcaacctatg ctgatagacc attaaataca gtttttgggt tcacagcttg tcatcatcat ttgtctatac 701 ctgtggcaaa gaatatctaa taagatactc tcagcatttt gcacacttaa actaagatgc 761 tgaatgctgt attttacgga ataatcagcc acattaaatt tggagactca acaagcatgc 821 tgtgaacatt caacattagg tttaaatttt atttttaaaa gttaataata aaaggatata 881 941 tgttaagtat tatgaaaccc tgcatatact gtaataaaat ggtggatgtg aatggacaat atatgcaata aaatttataa tttgattcya aaaaaaaaaa aamccv

```
<210> 340
```

<211> 748

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 372..494

<221> sig_peptide

<222> 372..443

<223> Von Heijne matrix

score 5.30000019073486

seg RILLLHFYCLLRS/SE <221> polyA_signal <222> 708..713 <221> polyA site <222> 732..745 <400> 340 acatgaaatg tgcttggtct gtgatctctt ggtcagatat ctgccttcca ggcgatcctt tgaggttgtg taattcagct ggccctggct cctggtccct gttactgagc tgggcagtcg 120 aaccgaaggc agatgagctc aagatcatgc cttgggaagc atggtgctct aggggtgcct 180 ttttattcct ttcattgtat tatagactgt ttccaagttt atggttagaa atggtaaagt 240 gggtctggtg ttttgaggta gaacccagcc tagggcaaga tatgaactgt tcttgaggta 300 gaaatgtcta cagtcagttg tttcatctag cttgcatctt aaaacacaaa cccttcagtt 360 410 gctttcactt a atg cac aca ttt gcc aat gac aga ggg tta tac agg atc Met His Thr Phe Ala Asn Asp Arg Gly Leu Tyr Arg Ile -15 -20 458 ctt ctt tta cat ttc tat tgt ctg cta cgc tca tca gag tat att ttg Leu Leu Leu His Phe Tyr Cys Leu Leu Arg Ser Ser Glu Tyr Ile Leu -10 ggg tac aag gtt ttg ggg gtt ttt tty ccc att ttg taactgeett 504 Gly Tyr Lys Val Leu Gly Val Phe Phe Pro Ile Leu 15 10 attgaaaadt aaktgccctt ccattccagg cctcctcata ttgtacttgt ttcctgccaa atctggggga tcatttgtat tttaactttg taatctatgg ctctgtactg ttgaaagstc 624 tcaattctgt ggggtctcct tagtatgtat gtgacttttc atgttgcaat atcacacgat 684 gggatggccc gacttttgct cttaataaat aatctgaatg agtaagaraa aaaaaaaaaa 744 748 <210> 341 <211> 1106

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 112..450

<221> sig_peptide

<222> 112..192

<223> Von Heijne matrix score 7.19999980926514 seq SLLFFLLLEGGXT/EQ

<221> polyA_signal

<222> 1053..1058

<221> polyA_site

<222> 1095..1106

<400> 341

aagac	ctcgg aa	acgagago cnnnnrct	g ccccgg t gaaggt	ggag cto	eggagege accttttg	gtgcacgcg	ca a ato	vacgga g aga 1 : Arg	60 .17
raa a	ag tgg a	aaa atq	qqa qqc	atq aaa	tac atc	ttt tcg t	tg ttg	ttc 1	165
Xaa L	ys Tro I	Lys Met	Gly Gly	Met Lys	Tyr Ile	Phe Ser I	Leu Leu	Phe	
-25	-	-	-20	_	-15			-10	
ttt c	tt ttg	cta gaa	gga ggc	kaa aca	gag caa	gtr amn o	cat tca	gag 2	213

Phe Leu Leu	Leu Glu Gl -5	y Gly Xaa	Thr Glu	Gln Val	Xaa His 5	Ser Glu	
aca tat tgo Thr Tyr Cys	atg ttt ca Met Phe Gl	a gac aag n Asp Lys 15	aag tac Lys Tyr	aga gtg Arg Val	ggt gag Gly Glu 20	aga tgg Arg Trp	261
cat cct tac His Pro Tyr 25	ctg gaa co Leu Glu Pr	t tat ggg to Tyr Gly 30	ttg gtt Leu Val	tac tgc Tyr Cys 35	gtg aac Val Asn	tgc atc Cys Ile	309
tgc tca gag	g aat ggg aa n Asn Gly As 45	n Val Leu	tgc agc Cys Ser	cga gtc Arg Val 50	aga tgt Arg Cys	cca aat Pro Asn 55	357
gtt cat tg	ctt tot co Leu Ser Pi	et gtg cat	att cct Ile Pro 65	cat ctg His Leu	tgc tgc Cys Cys	cct cgc Pro Arg 70	405
tgc cca gaa Cys Pro Gli	a gac tcc tt 1 Asp Ser Le	a ccc cca eu Pro Pro	gtg aac Val Asn 80	aat rwg Asn Xaa	gtg acc Val Thr 85	agc Ser	450
	<pre>\75 agtacaatgg</pre>	ananatta.	• -	rass add	tattcat	agctgrrggg	510
tagtettgek	atcggcaacc	gacaactta	c acceat	taca act	gttcgga	ragaaacktg	570
ctctttcaga	tcaagacttg	caccasatt	a acctot	acct tee	cagtete	tottccarat	630
tattgtggtc	gggtwtgcag	argagatgg	a caacto	tcat ggg	aacmttc	tgatggtgat	690
teetgetgee	aacctgccaa	cadadaadc	a agacati	tott acc	accactc	tcactatgat	750
acticagge	gccgacaggc	tagagaaga	a tecede.	tttc ctq	gggccag	aagtcaccgg	810
ggaggtgtta	tggattccca	gcaagcatc	a ddaacc	atta tac	aaattgt	catcaataac	870
ggageceeca	atggacaagt	aratattte	c aatgga	aaga cct	attotca	tggcgagtcc	930
tagcacccaa	acctccgggc	atttggcat	t atagaa	tata tac	tatgtac	ttgtaatgtc	990
accaadcaad	agtgtaagaa	aatccactq	c cccaat	cgat acc	cctgcaa	gtatcctcaa	1050
aaaatagacg	gaaaatgctg	caaggtgtg	t ccaggt	aaaa aag	caaaaaa	aaaaaa	1106

<400> 342 aaaacccagc ctacctgctg tagctgccgc cactgccgtc tagagcbnmag ccccagagcc taggaacctg gggcccgctc taggaacctg gggcccgct taggaacctg ggaacctg gggcccgct taggaacctg gggcccgct taggaacctg ggaacctg gaacctg ggaacctg gaacctg	teegeegeea etggweeeee 60 eteeceete eaggee atg 119 Met
agg att ctg cag tta atc ctg ctt gct ctg gca a Arg Ile Leu Gln Leu Ile Leu Leu Ala Leu Ala deu	aca ggg ctt gta ggg 167 Thr Gly Leu Val Gly -5
gga gag acc agg atc atc aag ggg ttc gag tgc gly Glu Thr Arg Ile Ile Lys Gly Phe Glu Cys 1	aag cct cac tcc cag 215 Lys Pro His Ser Gln 15

ccc Proʻ	tgg Trp	cag Gln	gca Ala	gcc Ala 20	ctg Leu	ttc Phe	gag Glu	aag Lys	acg Thr 25	cgg Arg	cta Leu	ctc Leu	tgt Cys	ggg 30	gcg Ala	263
Thr	Leu	Ile	Ala 35	ccc Pro	Arg	Trp	ctc Leu	Leu 40	Thr	Ala	Ala	nis	45	Deu	Dy B	311
Pro	Arg	Tyr	ata Ile	Xaa	His	Leu	999 55	Gln	His	Asn	Leu	60 61n	гÀг	GIU	GIU	359
Gly	Cys	gag Glu	Gln	Thr	Arg	Thr	gcc Ala	Thr	GIU	ser	75	PIO	UIS	PIO	Gly	407
Phe	Asn	Asn	Ser	Leu	Pro	Asn	aaa Lys	Asp	Xaa	Xaa 90	Asn	Asp	TIE	Mec	95	455
gtg Val	Xaa	Met	·Xaa	Ser	Pro	Val	tcc Ser	IIe	105	Trp	Ala	vai	Arg	110	neu	503
Thr	Leu	Ser	Ser	Arg	Cys	Val	act Thr	Ala 120	Gly	Thr	Ser	Cys	Leu 125	TIE	Ser	551
Gly	Trp	Gly	agc Ser	Thr	Ser	Ser	ccc Pro 135	Gln	Leu	Arg	Leu	140	HIS	TIIT	neu	599
Arg	Cys	gcc Ala	aac Asn	Ile	Thr	Ile 150	att Ile	Glu	His	Gln	Lys 155	Cys	GIU	Asn	Ala	647
tac Tyr 160	ccc Pro	aac	aac Asn	atc Ile	aca Thr 165	Asp	acc Thr	atg Met	gtg Val	tgt Cys 170	Ala	agc Ser	gtg Val	cag Gln	gaa Glu 175	695
aaa	aac	aag Lys	gac Asp	tcc Ser 180	tgc Cys	caq	ggt Gly	gac Asp	tcc Ser 185	GIA	ggc	cct	ctg Leu	gtc Val 190	tgt Cys	743
aac Asn	cag Gln	tct Ser	ctt Leu 195	caa Gln	aac	att Ile	atc :Ile	tcc Ser	Trp	ggc	cag Gln	gat Asp	ccg Pro 205	Cys	gcg Ala	791
atc Ile	acc	cga Arg	aag Lys	cat	ggt Gly	gtc Val	tac Tyr 215	acg Thr	aaa	gtc Val	tgo Cys	aaa Lys 220	ТУІ	gtg Val	gac	839
tgg Trp	Ile	caç Glr	gag	acg Thr	atg Met	aag Lys	aac Asn	aat	tag	actg	gac	ccac	ccac	ca		886
acc tca	ctaa ctta tatt	itca igcc iata	aaga atca acto	iccct iacct ctgg?	ct a gg g aa t	cact cgaa gttc	tggt catt gaaa acac	c tt at ca cc to	tggg gtga ggttt	geete igaee igtte	tgg tct	gact gatto g tt o	aca aaa tat	ttct	taagaa gatgotg goottg agooco waaaaaa	946 1006 1066 1126 1186
aaa		1900	رددو	guua	ica C		•~99					- د - ر		_		1191

<210> 343

<211> 1070

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 13..465

<221> sig_peptide <222> 13..75

<223> Von Heijne matrix
score 3.90000009536743
seq PVAVTAAVAPVLS/IN

<221> polyA_signal <222> 1035..1040

<221> polyA_site <222> 1060..1070

<400> 343 agagtcggga aa atg gct gcg agt acc tcc atg gtc ccg gtg gct gtg acg 51 Met Ala Ala Ser Thr Ser Met Val Pro Val Ala Val Thr -10 -15 -20 99 geg gea gtg geg eet gte etg tee ata aac age gat tte tea gat ttg Ala Ala Val Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu 1 **\-5** 147 egg gaa att aaa aag caa etg etg ett att geg gge ett ace egg gag Arg Glu Ile Lys Lys Gln Leu Leu Leu Ile Ala Gly Leu Thr Arg Glu . 20 15 cgg ggc cta cta cac agt agc aaa tgg tcg gcg gag ttg gct ttc tct 195 Arg Gly Leu Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser 35 3.0 25 ctc cct gca ttg cct ctg gcc gag ctg caa ccg cct ccg cct att aca 243 Leu Pro Ala Leu Pro Leu Ala Glu Leu Gln Pro Pro Pro Pro Ile Thr 50 45 gag gaa gat gee cag gat atg gat gee tat acc etg gee aag gee tae 291 Glu Glu Asp Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr 65 60 ttt gac gtt aaa gag tat gat cgg gca gca cat ttc ctg cat ggc tgc 339 Phe Asp Val Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys 80 aat gca aga aaa gcc tat ttt ctg tat atg tat tcc aga tat ctg gtg 387 Asn Ala Arg Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val 100 95 agg gcc att tta aaa tgt cat tct gcc ttt agt gaa aca tcc ata ttt 435 Arg Ala Ile Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe 120 115 110 aga acc aat gga aaa gtt aaa tct ttt aaa tagcttagca gtgggccact 485 Arg Thr Asn Gly Lys Val Lys Ser Phe Lys 130 125 gaatgaatgt actttataca tagcaataat aaaaaaaaga tatcataaat aaagttaaaa aggatggtaa aaaaaaaat attcttagga atgactaaca ggataagtaa caacctgatt atttatttac tttaggttat ataaggttct tcatgcctgt gaattaatat tattgtgtaa 665 gaattaagtt aaaaagcctg ggctgacttt taaatttata aattcattta tcatgtttat 725 agtatattta ttgtttttct ttcatggcta ttaaaaagta tgactgtaaa ggacaatgca 785 agtaaaccaa cttaatactg tattgaataa taagtacaat ttattattt actttgaaac attatgaatt tactttccta ctttttctta gttgttatct atataaattg attaaaaaaa cattttatgt acttctcatt tcctagtaca ggttgagtat cccttatttg aagtgcttgg 965 1025 gaccaaaagt gtttcagatt tcagattttt ttcagatttt ggtatatttg cattatactt 1070 actggttgaa ataaaaaatg ctgcagtgag tgtcaaaaaa aaaaa

<210> 344

<211> 1213

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 2..718

<221> sig_peptide <222> 2..76 <223> Von Heijne matrix score 3.90000009536743 seq RVGLLLGGGGVYG/SR <221> polyA_signal <222> 1170..1175 <221> polyA_site <222> 1203..1213 <400> 344 a atg ccc cgg aag cgg aag tgc gat ctt cgg gct gtc aga gtt ggt ctg 49 Met Pro Arg Lys Arg Lys Cys Asp Leu Arg Ala Val Arg Val Gly Leu -15 -20 tta ctc ggt ggt ggc gga gtc tac gga agc cgt ttt cgc ttc act ttt 97 Leu Leu Gly Gly Gly Val Tyr Gly Ser Arg Phe Arg Phe Thr Phe cct ggc tgt aga gcg ctt tcc ccc tgg cgg gtg aga vtg cag aga cga 145 Pro Gly Cys Arg Ala Leu Ser Pro Trp Arg Val Arg Xaa Gln Arg Arg 15 193 agg tgc gag atg agc act atg ttc gcg gac act ctc ctc atc gtt ttt Arg Cys Glu Met Ser Thr Met Phe Ala Asp Thr Leu Leu Ile Val Phe 30 ate tet gtg tge acg get etg etc gea gag gge ata ace tgg gte etg 241 Ile Ser Val Cys Thr Ala Leu Leu Ala Glu Gly Ile Thr Trp Val Leu 50 45 gtt tac agg aca gac aag tac aag aga ctg aag gca gaa gtg gaa aaa 289 Val Tyr Arg Thr Asp Lys Tyr Lys Arg Leu Lys Ala Glu Val Glu Lys 60 cag agt aaa aaa ttg gaa aag aag gaa aca ata aca gag tca gct 337 Gln Ser Lys Lys Leu Glu Lys Lys Lys Glu Thr Ile Thr Glu Ser Ala ggt cga caa cag aaa aar aaa ata gag aga cdd kaa kas amc ctg arg 385 Gly Arg Gln Gln Lys Lys Ile Glu Arg Xaa Xaa Xaa Leu Xaa 90 aat aac aac aga gat cta tca atg gtt cga atg aaa tcc atg ttt gct 433 Asn Asn Asn Arg Asp Leu Ser Met Val Arg Met Lys Ser Met Phe Ala 115 110 att ggc ttt tgt ttt act gcc cta atg gga atg ttc aat tcc ata ttt 481 Ile Gly Phe Cys Phe Thr Ala Leu Met Gly Met Phe Asn Ser Ile Phe 130 125 120 gat ggt aga gtg gtg gca aag ctt cct ttt acc cct ctt tct tas rtc 529 Asp Gly Arg Val Val Ala Lys Leu Pro Phe Thr Pro Leu Ser Xaa Xaa 145 sra gga ctg tct cat cga aat ctg ctg gga gat gac acc aca gac tgt Xaa Gly Leu Ser His Arg Asn Leu Leu Gly Asp Asp Thr Thr Asp Cys 160 155 tec ttc att ttc ctg taw att ctc tgt act atg tcg att cga cag aac 625 Ser Phe Ile Phe Leu Xaa Ile Leu Cys Thr Met Ser Ile Arg Gln Asn 180 175 673 att cag aag att ctc ggc ctt gcc cct tca cga gcc gcc acc aag cag Ile Gln Lys Ile Leu Gly Leu Ala Pro Ser Arg Ala Ala Thr Lys Gln 190 185 718 gca ggt gga ttt ctt ggc cca cca cct cct tct ggg aag ttc tct Ala Gly Gly Phe Leu Gly Pro Pro Pro Pro Ser Gly Lys Phe Ser 205 tgaactcaag aactctttat tttctakcat tctttctaga cacacacac tcagactggc 778 aactgttttg tascaagagc cataggtagc cttackactt gggcctcttt ctagttttga 838 898 attatttcta agccttttgg gtatkattag agtgaaaatg gcagccagca aacttgatag

tgcttttggt cctage tgtttatgta atgaa tgggaccgac tctca agatttagaa gaaaa ttttttcaag ccaaa atgtaaaaaa aaaaa	aaaca aatagcat aggca ctgtgtat attta gtttgttt tacat gacataar	cc ttcttgtttc gc cctgcaagtt aa cccttgtaac	ggctgtctat g	agcatttag 1018 ttgttgttt 1138
<210> 345 <211> 978 <212> DNA <213> Homo sapie	ns			
<220> <221> CDS <222> 86709				
<221> polyA_sign <222> 943948	nal			
<221> polyA_site <222> 963973	2			
<400> 345				atgcctcgg 60
aaagcatcct tccct ggacgaaaga gtcgg	redeed eegta at	tg cga gag ccg et Arg Glu Pro -90	g cag aag aga	acc gca 112
aca atc gca aaa Thr Ile Ala Lys -80	tyc rrg gcs to Xaa Xaa Ala Xa	va gag ggc cto	c cga gac ccc Arg Asp Pro -70	tat ggc 160 Tyr Gly
cgc ctc tgt ggt Arg Leu Cys Gly -65	Ser Glu His P	cc cga aga cca ro Arg Arg Pro 60	a cct gag cgg D Pro Glu Arg -55	ccc gag 208 Pro Glu
gaa gac ccg agc Glu Asp Pro Ser -50	act cca gag gag Thr Pro Glu G:	ag gcc tct acc lu Ala Ser Thi	acc cct gaa Thr Pro Glu -40	gaa gcc 256 Glu Ala
tcg agc act gcc Ser Ser Thr Ala	caa gca caa a Gln Ala Glr ₃ L	ag cct tca gto ys Pro Ser Va -2!	l Pro Arg Ser	aat ttt 304 Asn Phe -20
cag ggc acc aag Gln Gly Thr Lys	aaa agt ctc c Lys Ser Leu L	tg atg tct ata eu Met Ser Ilo -10	a tta gcg ctc e Leu Ala Leu	atc ttc 352 Ile Phe -5
atc atg ggc aac Ile Met Gly Asn 1	age gee aag g	lu Ala Leu Va	tgg aaa gtg l Trp Lys Val 10	ctg ggg 400 Leu Gly
aag tta gga atg Lys Leu Gly Met 15	Gln Pro Gly A 20	arg Xaa His Se	r Ile Phe Gly 25	Asp Pro
aag aar atc gtc Lys Lys Ile Val 30	Thr Glu Xaa P	he Val Arg Ar 40	g Gly Tyr Leu	Ile Tyr 45
ara ccg gtg ccc Xaa Pro Val Pro	cgt abe agt c Arg Xaa Ser P 50	ccg gtg gag ta Pro Val Glu Ty 55	t gas ttc ttc r Xaa Phe Phe	tgg ggg 544 Trp Gly 60

ccc cga gca cac gtg gaa tcg agc ara ctg aaa stc wtg cat ttt gtg Pro Arg Ala His Val Glu Ser Ser Xaa Leu Lys Xaa Xaa His Phe Val 65 70 75
gca agg gtt cgt aac cga tgc tct aaa gac tgg cct tgt aat tat gac Ala Arg Val Arg Asn Arg Cys Ser Lys Asp Trp Pro Cys Asn Tyr Asp 80 85 90
tgg gat tcg gac gat gat gca gag gtt gag gct atc ctc aat tca ggt Trp Asp Ser Asp Asp Asp Ala Glu Val Glu Ala Ile Leu Asn Ser Gly 95 100 105
gct arg ggt tat tcc gcc cct taagtaratc tgaggcagac ccttgggggt 739 Ala Xaa Gly Tyr Ser Ala Pro 110 115
gtaaaagaga gtcacaggta ccccaaggag tagatgccag ggtcctaagt tgaaaatgmt 799
gtcgattggg ggcgggggac actgtatttg atatttgtga tcagtgatca ttgttcaact 859 gcgaaataga gtgtttgctt ttgataatgg aaaattgtat tcgttttaaa attccgtttg 919
ttgagaataa caatatgttt aaaaatataa ttgaacaaat tttaaaaaaa aaaamcccy 978
<210> 346 <211> 810 <212> DNA
<213> Homo sapiens
<220> <221> CDS <222> 63320
<221> sig_peptide <222> 63179 <223> Von Heijne matrix score 3.90000009536743 seq VLAIGLLHIVLLS/IP
<221> polyA_signal
<221> polyA_site <222> 799810
<400> 346
agggaaccga tcccgggccg ttgatcttcg gccccacacg aacagcagag aggggcatca gg atg aat gtk ggc aca gcg cac ags dag gtg aac ccc aac acg cgg Met Asn Val Gly Thr Ala His Xaa Xaa Val Asn Pro Asn Thr Arg -35 -30 -25
gtk atg aac agc cgt ggc atc tgg ctc tcc tac gtg ctg gcc atc ggt Val Met Asn Ser Arg Gly Ile Trp Leu Ser Tyr Val Leu Ala Ile Gly -20 -15 -10
ctc ctc cac atc gtg ctg ctg agc atc ccg ttt gtk agt gtc cct gtc Leu Leu His Ile Val Leu Leu Ser Ile Pro Phe Val Ser Val Pro Val -5 1 5
gtc tgg acc ctc acc aac ctc att cac aac atg ggc atg tat atc ttc Val Trp Thr Leu Thr Asn Leu Ile His Asn Met Gly Met Tyr Ile Phe 10 20
ctg cac acg gtg aag ggg aca ccc ttt gag acc ccg gac cag ggc aag Leu His Thr Val Lys Gly Thr Pro Phe Glu Thr Pro Asp Gln Gly Lys 25 30 35 40
gcg agg ctg cta acc cac tgg tgagcagatg gattatgggg tccagttcac 350 Ala Arg Leu Leu Thr His Trp 45
ggcctctcgg aakttcttga ccatcacacc catcgtgctg tacttcctca ccagcttcta 410 cactaaktac raccaaatcc attttgtgct caacaccgtg tccctgatra gcgtgcttat 470

ccccaagctg ccccagctcc acggaktccg gatttttgga atcaataakt actgaaaktg cascccttc ccctgcccag ggtggcaggg gaggggtagg gtaaaaggca tktgctgcaa chctgaaaac araaaraara rscctctgga cactgccara ratgggggtt gagcctctgg cctaatttcc cccctgctt cccccagtag ccaacttgga gtagcttgta ytggggttgg ggtaggccc ctgggctctg accttttctg aattttttga tcttttcctt ttgctttttg aatararact ccatggagtt ggtcatggaa aaaaaaaaaa	530 590 650 710 770 810
<210> 347 <211> 771 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 299418	
<pre><221> sig_peptide <222> 299379 <223> Von Heijne matrix</pre>	
<221> polyA_signal <222> 739744	
<221> polyA_site <222> 762771	
<400> 347 accttgggct ccaaattcta gctcataaag atgcaagtkt tgcaatttcc tataaatggt taagaaaaga gcaagctgtc cagagagtga gaagtttgaa aagagaggtg cataagagag aaatgatgtc catttgagcc ccaccacgga ggttatgtgg tcccaaaagg aatgatggcc aagcaattaa tttttcctcc tagttcttag cttgcttctg cattgattgg ctttacacaa ctggcattta gtctgcatta cacaaataga cactaattta tttggaacaa gcagcaaa atg aga act tta ttt ggt gca gtc agg gct cca ttt agt tcc ctc act Met Arg Thr Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr -25 -20 -15	60 120 180 240 298 346
ctg ctt cta atc acc cct tct ccc agc cct ctt cta ttt gat aga ggt Leu Leu Leu Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly -10 -5 1 5	394
ctg tcc ctc aga tca gca atg tct tagcccctct cctctcttcc attccttcct Leu Ser Leu Arg Ser Ala Met Ser 10	448
gttggtactc atttcttcta acttttaata aacatttagg tataatacat tacagtaagt gctatttaga tacaaactta aaacatacta tatattttaa ggatctaaga atcctttara rrrggcacat gactgaagta cctcagctgc gcagcctgta accagttttt ttaatgtaaa agtaaraatg ccagccttaa cctabccctg carataaaag ctaactttta ttaataccag ccctgaataa tggcactaat ccacactctt ccttaragtg atgctggaaa aataaaatca ggggcttcag attaaaaaa aaa	508 568 628 688 748 771

<210> 348

<211> 409

<212> DNA

<213> Homo sapiens

<220>

....

<221> CDS

<222> 186..380

<pre><221> sig_peptide <222> 186233 <223> Von Heijne matrix score 4 seq FFLFLSFVLMYDG/LR</pre>	
<221> polyA_signal <222> 383388	
<221> polyA_site <222> 396409	
<400> 348 ataaaagaag cagcaaatag aatttcccac aaagtaagtt gactctaaat cttaagtatt acctagttt ttaaaggttt gaatataata atgcagtatt tgcagtataa aaaggaagga atttgtagag aatcattttg gtgctcaagt ctcttagcag tgccttattg cctcatagca agaag atg ctg ggg ttt ttt ttg ttt ttg tcc ttt gta tta atg tat gat Met Leu Gly Phe Phe Leu Phe Leu Ser Phe Val Leu Met Tyr Asp -15 -10 -5	60 120 180 230
ggt ttg cgc ctt ttt ggc att ctt tca aca tgt cgt gta cat cac acc Gly Leu Arg Leu Phe Gly Ile Leu Ser Thr Cys Arg Val His His Thr 1 5 10 15	278
atg aat cag ttc cta att gat ata tct agc ttt acc tcc cga gtt aaa Met Asn Gln Phe Leu Ile Asp Ile Ser Ser Phe Thr Ser Arg Val Lys 20 25 30	326
aaa aaa atc ttt tta ttt tat goc ttc awa ggt tgc ycg ttt car agt Lys Lys Ile Phe Leu Phe Tyr Ala Phe Xaa Gly Cys Xaa Phe Gln Ser	374
35 40 45 gcc aca taaataaaat gtttaacaaa aaaaaaaaa Ala Thr	409
<pre><210> 349 <211> 613 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 69458 <221> sig_peptide <222> 69233 <223> Von Heijne matrix</pre>	
aagaacctga gcagcctgtc ttcagacaga gagaggccca cggctgtttc ttgaaaytgg cgctggga atg gcc atg tgg aac agg cca tgb bag ang ctg cct cag cag Met Ala Met Trp Asn Arg Pro Xaa Xaa Xaa Leu Pro Gln Gln -55 -50 -45	60 110
cct cts sta gct gag ccc act gca gag ggg gag cca cac ctg ccc acg Pro Leu Xaa Ala Glu Pro Thr Ala Glu Gly Glu Pro His Leu Pro Thr	158

-40 -35 -30	
ggc cgg gas byg act gag gcc aac cgc ttc gcc tat gct gcc ctc tgt Gly Arg Xaa Xaa Thr Glu Ala Asn Arg Phe Ala Tyr Ala Ala Leu Cys -25 -20 -15 -10	206
ggc atc tcc ctg tcc cag tta ttt cct gaa ccc gaa cac agc tcc ttc Gly Ile Ser Leu Ser Gln Leu Phe Pro Glu Pro Glu His Ser Ser Phe -5 1 5	254
tgc aca gag ttc atg gca ggc ctg gtg ckm tgg ctg gag ttg tct gaa Cys Thr Glu Phe Met Ala Gly Leu Val Xaa Trp Leu Glu Leu Ser Glu 10 15 20	302
gct gtc ttg cca acc atg act gct ttt gcg agc ggc ctg gga ggt gaa Ala Val Leu Pro Thr Met Thr Ala Phe Ala Ser Gly Leu Gly Gly Glu 25 30 35	350
gga sca vma tgt gtt tgt tca aat ttt act gaa gga ccc cat ctt gaa Gly Xaa Xaa Cys Val Cys Ser Asn Phe Thr Glu Gly Pro His Leu Glu 40 45 50 55	398
gga cga ccc gac ggt gat cac tca gga cct tct gag ctt ctc act caa Gly Arg Pro Asp Gly Asp His Ser Gly Pro Ser Glu Leu Leu Thr Gln 60 65 70	446
gga tgg gca cta tgacscccgg gccagagtcc tcgtttgcca catgacctcc Gly Trp Ala Leu 75	498
ctgctccaag tgcccttgga ggagctggat gtccttgaaa agatgttcct ggagagcctg aaggaaatca aagaagagga atctgaaatg gccgaggcat cccraaaaaa aaaaa	558 613
<pre><210> 350 <211> 986 <212> DNA <213> Homo sapiens </pre> <pre><220> <221> CDS <222> 12638 </pre> <pre><221> sig_peptide <222> 12263 </pre> <pre><223> Von Heijne matrix</pre>	
accetateaa g atg gte aac tte eee cag aaa att gea ggt gaa ete tat Met Val Asn Phe Pro Gln Lys Ile Ala Gly Glu Leu Tyr -80 -75	50
gga cct ctc atg ctg gtc ttc act ctg gtt gct atc cta ctc cat ggg Gly Pro Leu Met Leu Val Phe Thr Leu Val Ala Ile Leu Leu His Gly -70 -65 -60	98
atg aag acg tct gac act att atc cgg gag ggc acc ctg atg ggc aca Met Lys Thr Ser Asp Thr Ile Ile Arg Glu Gly Thr Leu Met Gly Thr -55 -40	146
gcc att ggc acc tgc ttc ggc tac tgg ctg gga gtc tca tcc ttc att Ala Ile Gly Thr Cys Phe Gly Tyr Trp Leu Gly Val Ser Ser Phe Ile -35 -30 -25	194
tac ttc ctt gcc tac ctg tgc aac gcc cag atc acc atg ctg cag atg	242

			-20					-15		Ile			-10		· ·	
Leu	Ala	Leu	Leu	Gly	Tyr	Gly	Leu 1	Phe	Gly	Cat	Cys 5	11e	vai	Den	FILE	290
Ile	Thr	tat Tyr	Asn	Ile	His 15	Leu	Arg	Ala	Leu	ttc Phe 20	Tyr	Leu	Pne	ттр	25	338
tta	gtg Val	ggt Gly	gga Gly	ctg Leu 30	tcc Ser	aca Thr	ctg Leu	cgc Arg	atg Met 35	gta Val	gca Ala	gtg Val	ttg Leu	gtg Val 40	tct Ser	386
cgg Arg	acc Thr	gtg Val	ggc Gly 45	ccc	aca Thr	cad Xaa	cgg Arg	mtg Xaa 50	ctc Leu	ctc Leu	tgt Cys	ggc	acc Thr 55	ctg Leu	gct Ala	434
gcc Ala	cta Leu	cac His 60	atg	ctc Leu	ttc Phe	ctg Leu	ctc Leu 65	tat Tyr	ctg Leu	cat His	ttt Phe	gcc Ala 70	tac Tyr	cac His	aaa Lys	482
dtg Xaa	gta Val 75	dag	gjå aaa	atc Ile	ctg Leu	gac Asp 80	aca Thr	ctg Leu	gag Glu	ggc Gly	ccc Pro 85	aac Asn	atc Ile	ccg Pro	ccc Pro	530
atc Ile 90	cag	agg Arg	gtc Val	ccc Pro	aga Arg 95	gac Asp	atc Ile	cct Pro	gcc Ala	atg Met 100	ctc Leu	cct Pro	gct Ala	gct Ala	cgg Arg 105	578
ctt	ccc Pro	acc Thr	acc Thr	gtc Val 110	ctc Leu	aac Asn	gcc Ala	aca Thr	gcc Ala	Lys	gct Ala	gtt Val	gcg Ala	gtg Val 120	acc Thr	626
			cac His	tga		acc	tgaa	atto		gcca	gtcc	t ct	ttcc	cgca		678
ttt tga aaa ccc	gcag aagg tggg	ctg cac tca tgt	ggar ccac aagg	gaas tgag ccaa cttt	ct g ga a ga g	tago ctco aaco	tgcg tggc cctc	t aa c ag	igtac gact acct	ctcc gcaa accc	ttg ggc ctt	atgo tctg cctt	cag	ccaa cttt	atgggg cacttc tgcaga atctct ggaaaa	738 798 858 918 978 986

<210> 351

<211> 1447

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 282..389

<221> sig_peptide

<222> 282..332

<223> Von Heijne matrix score 3.5 seq RWWCFHLQAEASA/HP

<221> polyA_signal

<222> 1413..1418

<221> polyA_site

<222> 1437..1447

<400> 351
ataataatat ctaaaaagct aaattttaaa taccagcttt acataaatga ttgtkgactc
tggtctgtkt ctgacacctt tccagaaaaa agtcaattgt tcaggtacac caaagaggaa 120

gaagagetgt ggaggeeace etetacaaag etttatagaa ettetggate taacteacaa acaagettee agaagagaet agagacetta ggeeaggaga tgaaggagtt eagtageaaa gteacacetg tecaatteee tgagetttge teacteaget a atg gga tgg caa agg Met Gly Trp Gln Arg -15	180 240 296
tgg tgg tgc ttt cat ctt cag gca gaa gcc tct gcc cat ccc cct caa Trp Trp Cys Phe His Leu Gln Ala Glu Ala Ser Ala His Pro Pro Gln -10 -5 1	344
ggg ctg cag gcc caa ttc tca tgc tgc cct tgg gtg ggc atc tgt Gly Leu Gln Ala Gln Phe Ser Cys Cys Pro Trp Val Gly Ile Cys 5 10 15	389
taacaaadga aaacgtctgg gtggcggcag casctttgct ctgagtgcct acaaagctaa	449
tgcttggtgc tagaaacatc atcattatta aacttcagaa aagcagcagc catgttcagt	509 569
caggeteatg etgeeteact gettaagtge etgeaggage egeetgeeaa reteceette	629
ctacacctgg cacactgggg tetgcacaag getttgtcaa ccaaaracag ettececeww ttgattgeet gtagaetttg gagecaaraa acactetgtg tgaetetaca cacaetteag	689
gtggtttgtg cttcaaagtc attgatgcaa cttgaaagga aacagtttaa tggtggaaat	749
gaactaccat ttataacttc tgttttttta ttgagaaaat gattcacgaa kkccaaatca	809
gattgccagg aagaaatagg acgtgacggt actgggccct gtgattctcc cagcccttgc	869
agtccgctag gtgagaggaa aagctcttta cttccgcccc tggcagggac ttctgggtta	929
tqqqaqaaac caqagatggg aatgaggaaa atatgaacta cagcagaagc ccctgggcag	989
ctgtgatgga gcccctgaca ttactcttct tgcatctgtc ctgccttctt tccctctgcg	1049 1109
aggcagtggg gtgggattca gagtgcttag tctgctcact gggagaagaa gagttcctgc	1169
gcatgcaago cotgotgtgt ggctgtcgtt tacatttggg aggtgtcctg tatgtctgta cgttggggac tgcctgtatt tggaagattt aaaaacctag catcctgttc tcaccetcta	1229
agotgoattg agaaatgact ogtototgta tttgtattaa goottaacac ttttottaag	1289
tgcattcggt gccaacattt tttagagctg taccaaaaca aaaagcctgt actcacatca	1349
camtgtcatt ttgataggag cgttttgtta tttttacaag gcagaatggg gtgtaacagt	1409
tgaattaaac ttagcaatca cgtgctcaaa aaaaaaaa	1447
<pre><210> 352 <211> 1641 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 208339 <221> sig_peptide <222> 208294 <223> Von Heijne matrix</pre>	
<400> 352 agaaccgtga tgggaagatg gacaaggaag agaccaaaga ctggatcctt ccctcagact atgatcatgc agaggcagaa gccaggcacc tggtctatga atcagaccaa aacaaggatg gcaagcttac caaggaggag atcgttgaca agtatgactt atttgttggc agccaggcca cagattttgg ggaggcctta gtacggc atg atg agt tct gag cta cgg agg aac Met Met Ser Ser Glu Leu Arg Arg Asn -25	60 120 180 234
cct cat ttc ctc aaa agt aat tta ttt tta cag ctt ctg gtt tca cat Pro His Phe Leu Lys Ser Asn Leu Phe Leu Gln Leu Leu Val Ser His -20 -15 -10 -5	282
gaa att gtt tgc gct act gag act gtt act aca aac ttt tta aga cat Glu Ile Val Cys Ala Thr Glu Thr Val Thr Thr Asn Phe Leu Arg His	330

```
gaa aag gcg taatgaaaac catcccgtcc ccattcctcc tcctctctga
                                                                      379
        15
gggactggag ggaagccgtg cttctgagga acaactctaa ttagtacact tgtgtttgta
                                                                      439
                                                                      499
ratttacacw wtgtattatg tattaacatg gcgtgtttat ttttgtattt ttctctggtt
                                                                      559
gggagtatka tatgaaggat caarateete aacteacaca tgtaracaaa cattasetet
                                                                      619
ttactctttc tcaacccctt wtatgatttt aataattctc acttaactaa ttttgtaagc
                                                                      679
ctgagatcaa taagaaatgt tcaggagaga ggaaagaaaa aaaatatatg ctccacaatt
                                                                      739
tatatttaga gagagaacac ttagtcttgc ctgtcaaaaa gtccaacatt tcataggtag
taggggccac atattacatt cagttgctat aggtccagca actgaacctg ccattacctg
                                                                      799
ggcaaggaaa gatccctttg ctctaggaaa gcttggccca aattgatttt cttcttttc
                                                                      859
cccctgtagg actgactgtt ggctaatttt gtcaagcaca gctgtggtgg gaagagttag
                                                                      919
ggccagtgtc ttgaaaatca atcaagtagt gaatgtgatc tctttgcara gctatagata
                                                                     979
gaaacagctg gaaaactaaa ggaaaaatac aagtgttttc ggggcataca ttttttttct
                                                                     1039
gggtgtgcat ctgttgaaat gctcaagact taattatttg ccttttgaaa tcactgtaaa
tgcccccatc cggttcctct tcttcccarg tgtgccaagg aattaatctt ggtttcacta
                                                                     1159
caattaaaat toactoottt coaatcatgt cattgaaagt gootttaacg aaagaaatgg
                                                                     1219
tcactgaatg ggaattctct taagaaaccc tgagattaaa aaaagactat ttggataact
tataggaaag cctagaacct cccagtagag tggggatttt tttcttcttc cctttctctt
ttggacaata gttaaattag cagtattagt tatgagtttg gttgcagtgt tcttatcttg
                                                                    1399
tgggctgatt tccaaaaacc acatgctgct gaatttacca gggatcctca tacctcacaa
                                                                     1459
                                                                   1519
tgcaaaccac ttactaccag gcctttttct gtgtccactg gagagcttga gctcacactc
aaagatcaga ggacctacag agagggctct ttggtttgag gaccatggct tacctttcct
                                                                   1639
goottigaco catcacacco catticotoc totticocto teccogotgo caaaaaaaaa
                                                                     1641
aa
<210> 353
<211> 884
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 69..557
<221> sig_peptide
<222> 69..224
<223> Von Heijne matrix
      score 4.69999980926514
      seq LGLALGRLEGGSA/RH
<221> polyA_signal
<222> 849..854
<221> polyA_site
<222> 870..883
<400> 353
attggctccg gatcgtgcgt gaggcggctt cgtgggcagc gagagtcaca gacaagacag
                                                                      60
caagcagg atg gag cac tac cgg aaa gct ggc tct gta gag ctc cca gcg
                                                                      110
         Met Glu His Tyr Arg Lys Ala Gly Ser Val Glu Leu Pro Ala
                 -50
                                     -45
cct tcc cca atg ccc cag cta cct cct gat acc ctt gag atg cgg gtc
                                                                      158 .
Pro Ser Pro Met Pro Gln Leu Pro Pro Asp Thr Leu Glu Met Arg Val
                                -30
cga gat ggc agc aaa att cgc aac ctg ctg ggg ttg gct ctg ggt cgg
                                                                     206
Arg Asp Gly Ser Lys Ile Arg Asn Leu Leu Gly Leu Ala Leu Gly Arg
                            -15
                                                -10
ttg gag ggc ggc agt gct cgg cat gta gtg ttc tca ggt tct ggc agg
                                                                     254
```

	•	•															
L	eu		Gly	Gly	Ser	Ala		His	Val	Val	Phe	Ser	Gly	Ser	Gly	Arg 10	•
		-5		~	~~+	gtc	1	tac	act	gag	5 att	atc	aaq	caa	cgg		302
9	ICT.	gca	gga	Lve	Δla	Val	Ser	Cvs	Ala	Glu	Ile	Val	Lys	Arg	Arg	Val	
					15					20					25		
c	cq.	ggc	ctg	cac	cag	ctc	acc	aag	cta	ckt	ttc	ctt	caa	act	gag	gac	350
F	ro	Gly	Leu	His	Ġln	Leu	Thr	Lys	Leu	Xaa	Phe	Leu	Gln	Tiir	Glu	Asp	
				30					35					40		ata	398
ā	gc	tgg	gtc	cca	scc	tca	cct	gac	aca	999	cta	rac	Dro	Tien	Thr	Val	J J G
٤	er	Trp		Pro	Xaa	Ser	Pro	Asp 50	THE	GIY	Leu	Aaa	55	200		,	
_			45	~+~	cct	gca	kta		ata	cta	ctc	asc		gac	ccc	ctg	446
7	gc	Ara	His	Val	Pro	Ala	Xaa	Trp	Val	Leu	Leu	Xaa	Arg	Asp	Pro	Leu	
		60					65					70					
c	gac	~~~	aat	gag	tgt	ggt	tac	caa	CCC	cca	gga	gca	CCC	cct	ggc	ctg	494
7	Asp	Pro	Asn	Glu	Cys	Gly	Tyr	Gln	Pro	Pro	GTÀ	Ala	Pro	Pro	GIY	Leu 90	
-	75					80					85		ara	222	agg		542
9	ggt	tcc	atg	CCC	agc	tcc	agc	tgt	ggc	Dro	Ara	Ser	Yaa	Livs	Ara	Ala	
(31y	Ser	Met	Pro	ser 95	ser	Ser	Cys	СТУ	100	Arg	501	11.44	-1-	105	Ala	
		rac	300	caa	75 75	taa	aaac	cta	ctga		qc c	tgtt	ctcc	g gg	cctr	aatg	597
				Arg		-5-		J			_	_					
				110													657
1	tct	9999	tgc	ttgt	gcct	tt t	ctra	naag	c gt	tgtg	askg	ctc	aaca	tcc	ccat	caaggt	657 717
	ttg	agtc	cac	aaaa	gtgg	ac c	taca	tato	a tg	cttc	ccct	tcc	ctct	agc	acgi	gggaag	777
9	gga	ctgc	tgt	gaag	aatg	ac a	gatg	tggg	g cc	cetg	tacc	ttc	tgaa	acc	aatc	aataag cakgga	837
	ggc'	ttcc -+++	tct	gcct	ecta 	tt t	ctac	ctat	t ta	aaaa	aaaa	aaa	acac			, , , , ,	884
	999	alll	agg	aaac	aaay			0000									
												- •					
	<21	0 > 3	54														
		1> 7															
		2> D		2													
	<21	3> H	omo.	sapi	ens												
	<22	0 >															
		1> C	DS														
			34	325													
				epti	.de												
			34														
	<22					trix 00095		1 2									
		_				NTHO		2 3									
		-	seq .		1001	2112111	-,										
	<22	21> r	olyA	A sit	:e												
			718.														
	<40	00> 3	354							++	~ = + <i>~</i> ·	-, <u>+</u> + + +	- 2012 1	agea	taa	cctataa	60
	ato	atti	tct	tate	cct	got 9	Jacci	taat	בט כי	taac	atta	a cti	tooti	aaa	taai	cctgtaa taatcaa	120
	tgt	aato	gcaa	tet	ato	cat	aat	ttt	gaa	ata	ata	tcc	ttg	aaa	gag	gaa	169
	-96	-uay			Met	His	Gly	Phe	Glu	Ile	Ile	Ser	Leu	Lys	Glu	Glu	
							-45					-40					
	tca	a cc	a tta	a gg	a aa	g gt	g ag	t ca	g gg	t cc	t tt	g tt	t aat	t gt	g ac	t agt	217
	Sea	r Pr	o Le	u Gl	y Ly	s Va	l Se	r Gl	n Gl	y Pr	o Le	u Pho	e Ası	n Val	ı Th	r Ser -20	
	-35	5				- 3				~ ~~	-2.		c + c	- tt	c ca		265
	gge	c tc	a to	a to	a cc.	a gt	g ac I Th	ט נקי די די	y tt	y 99 u G)	v Le	u Le	u Se:	r Ph	e Gl	g aac n Asn	
					-1	5				-1	0				- 5		
	cto	q ca	t ta	c tt	c cc	- aga	c ct	c cc	c ac	t ga	g at	g cc	t ct	a ar	a gc	c aaa	313
		_	_			_											

1 5 10	
gga ktc aac act tgagcctagg gtgggctaca acaaaaratt ctaatttacc Gly Xaa Asn Thr	365
ttgcttcatc taggtccagg ccccaaktag cttgctgaag gaacttaaaa agtagctgtt atttattgta ttgtataasc taaaacatt tatttttgtt gaatcraaac aattccatgt ascaatcttt tttctgttca cggtgtttgt gataaaacct taaattccgc aagcatcagt tttttgaaaa aatgggaatt gaccggatag wwacaggcaa agwtataaat agctacaaca tcatttaact tttataaaca tgccttctct ctattgaara catctgatat ttttgctgga aagttggatc tatcctcagt aactctgcca tgaattcctg tttcckggtt ccaaaaaaaaa aaaa	425 485 545 605 665 725 729
<210> 355	
<211> 1013 <212> DNA	
<213> Homo sapiens	
<220> <221> CDS	
<222> 78731	
<221> sig_peptide <222> 78227	
<pre><223> Von Heijne matrix score 5.09999990463257</pre>	
seq RTALILAVCCGSA/SI	
<221> polyA_site <222> 10021013	
<400> 355	60
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Gly	60 110
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa	
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys -35 -30 -25	110
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu	110
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 -10 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa	110
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln -5	110 158 206 254
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 -10 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln	110 158 206
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln -5 aac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag Asn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys 10 25	110 158 206 254
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln -5 aac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag Asn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys 10 15 20 25 tat gct gtt tct agt cgt cat aat gta att tgc cag tta ctt tct gac Tyr Ala Val Ser Ser Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp	110 158 206 254 302
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aatttattt acttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln -5 aac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag Asn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys 10 15 20 25 tat gct gtt tct agt cgt cat aat gta att tgc cag tta ctt tct gac Tyr Ala Val Ser Ser Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp 30 35 40 tac aaa raa aaa cag atr cta aaa gtc tct tct tct gaa aac agc aat cca	110 158 206 254 302
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt acttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 -10 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln -5 ac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag Asn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys 10	110 158 206 254 302 350
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 -10 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln -5 aac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag Asn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys 10 15 20 25 tat gct gtt tct agt cgt cat aat gta att tgc cag tta ctt tct gac Tyr Ala Val Ser Ser Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp 30 35 tac aaa raa aaa cag atr cta aaa gtc tct tct tct gaa aac agc aat cca Tyr Lys Xaa Lys Gln Xaa Leu Lys Val Ser Ser Glu Asn Ser Asn Pro 45 55 raa caa gac tta aag ctg aca tca gag gaa gag tca caa agg ctt aaa Xaa Gln Asp Leu Lys Leu Thr Ser Glu Glu Glu Ser Gln Arg Leu Lys	110 158 206 254 302
agtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta aattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt Met His His Gly Leu Thr Pro Leu Leu Leu Gly -50 -45 -40 gta cat gag caa aaa cag caa gtg gtg aaaa ttt tta atc aag aaa aaa Val His Glu Gln Lys Gln Gln Val Lys Phe Leu Ile Lys Lys Lys -35 -30 -25 gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu -20 -15 gct gta tgt tgt gga tcg gca agt ata gtc agc ctt cta ctt gag caa Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln -5 aac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag Asn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys 10 15 20 25 tat gct gtt tct agt cgt cat aat gta att tgc cag tta ctt tct gac Tyr Ala Val Ser Ser Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp 30 35 40 tac aaa raa aaa cag atr cta aaa gtc tct tct ct gaa aac agc aat cca Tyr Lys Xaa Lys Gln Xaa Leu Lys Val Ser Ser Glu Asn Ser Asn Pro 45 raa caa gac tta aag ctg aca tca gag gaa gag tca caa agg ctt aaa	110 158 206 254 302 350

aat arg ggt ggt gat aga aag gtt gaa raa raa atg aar aag cac gga	542
Asn Xaa Gly Gly Asp Arg Lys Val Glu Xaa Xaa Met Lys Lys His Gly	
90 95 100 105	
agt wet cat atg gga tte cea raa aac etg met aac ggt gee act get	590
Ser Xaa His Met Gly Phe Pro Xaa Asn Leu Xaa Asn Gly Ala Thr Ala	
110 115 120	
gac aat ggt gat gga tta att ccm cca rgg aaa asc ara aca cct	638
Asp Asn Gly Asp Asp Gly Leu Ile Pro Pro Xaa Lys Xaa Xaa Thr Pro	
125 130 135	
qaa agc cas caa ttt cct gac act gag aat gaa cag tat cac agg gac	686
Glu Ser Xaa Gln Phe Pro Asp Thr Glu Asn Glu Gln Tyr His Arg Asp	
140 145 150	
ttt tot ggo cat ecc mac ttt ecc aed acc ett ecc atc aaa eag	731
Phe Ser Gly His Pro Xaa Phe Pro Thr Thr Leu Pro Ile Lys Gln	
155 160 165	
tgatgaacaa aatgatactc hsaagcmmct ttctgaagam caraacactg gaatattaca	791
agatgagatt ctgattcatg aagaaaagca gatagaagtg gctgaaaatg aattctgagc	851
tttctcttag ttataaraaa gaaaaagacc tcttgcatga aaatagtacg ttgcaggaag	911
	971
aaattgtcat gctaaractg gaactagack taatgaaaca tcagagccag ctaararaaa	
araaatattt ggaggaaatt gaaagtgtgg aaaaaaaaa	1013
<210> 356	
2207 330	

<211> 973

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 46..693

<221> sig_peptide

<222> 46..90

<223> Von Heijne matrix score 7.59999990463257 seq CVLVLAAAAGAVA/VF

<221> polyA_signal

<222> 937..942

<221> polyA_site <222> 962..973

<400> 356

<pre>aagcggctgg tccccggaag ttggacgcat gcgccgtttc tctgc atg gtg tgc gtt</pre>	57
ctc gtt cta gct gcg gcc gca gga gct gtg gcg gtt ttc cta atc ctg Leu Val Leu Ala Ala Ala Ala Gly Ala Val Ala Val Phe Leu Ile Leu -10 -5 1 5	105
cga ata tgg gta gtg ctt cgt tcc atg gac gtt acg ccc cgg gag tct Arg Ile Trp Val Val Leu Arg Ser Met Asp Val Thr Pro Arg Glu Ser 10 15 20	153
ctc agt atc ttg gta gtg gct ggg tcc ggt ggg cat acc act gag atc Leu Ser Ile Leu Val Val Ala Gly Ser Gly Gly His Thr Thr Glu Ile 25 30 35	201
ctg agg ctg ctt ggg agc ttg tcc aat gcc tac tca cct aga cat tat Leu Arg Leu Leu Gly Ser Leu Ser Asn Ala Tyr Ser Pro Arg His Tyr 40 45 50	249
gtc att gc t gac act gat gaa atg agt gcc aat aaa ata aat tct ttt Val Ile Ala Asp Thr Asp Glu Met Ser Ala Asn Lys Ile Asn Ser Phe	297

•																
	55					60					65					
gaa Glu	cta Leu	rat Xaa	cga Arg	gsk Xaa	gat Asp	aga Arg	rac Xaa	cct Pro	agt Ser	aac Asn	atg Met	twt Xaa	acc Thr	aaa Lys	tac Tyr	345
70					75					80					85	
tac Tyr	att Ile	cac His	cga Arg	Ile	cca Pro	ara Xaa	agc Ser	cgg Arg	gag Glu 95	gtt Val	cag Gln	cag Gln	tcc Ser	tgg Trp 100	Pro	393
tcc	200	att	two	90	200	tta	cac	tcc	-	taa	ctc	tcc	ttk	CCC	cta	441
														Pro		-
att Ile	cac His	agg Arg 120	gtg Val	aag Lys	cca Pro	rat Xaa	ttg Leu 125	gtg Val	ttg Leu	tgt Cys	aac Asn	gga Gly 130	cca Pro	gga Gly	aca Thr	489
tgt Cys	Val	cct	atc Ile	tgt Cys	gta Val	Ser	gcc	ctt Leu	ctc Leu	ctt Leu	Gly	ata	cta Leu	gga Gly	ata Ile	537
	135					140					145					r 0.5
aag Lys 150	aaa Lys	gtg Val	atc Ile	att Ile	yal 155	tac Tyr	gtt Val	gaa Glu	agc Ser	Ile 160	tgc Cys	Arg	yal Val	aaa Lys	Thr 165	585
	taa	ato	tac	gga		att	cta	ttt	cat		tca	aat	tac	ttc		633
														Phe 180		
														tac		681
Val	Gln	Trp	Pro 185	Ala	Leu	Lys	Glu	Lys 190	Tyr	Pro	Lys	Ser	Val 195	Tyr	Leu	
		att Ile 200		tgad	caaat	gg d	caact	zgact	it ct	ttag	gaatt	t ttg	gcast	taa		733
cagt	arta		acto	caaat	t gg	9999	gaaaa	a aaa	accct	caca	tgtt	tctt	gt a	aaagg	gcgtct	793
gaca	igtco	tg a	araat	tati	g at	ggta	aagga	a ata	aaaa	aatg	twc	agati	cac t	cagt	gaara	853
aact	gagg	gct t	ctct	tate	ga aa	acaaa	acatt	: gat	caaa	gta	acta	acyaa	aat 9	gttta	atgcct	913
ctgt	aaac	cca a	aattt	ctt	it ct	carat	caaaa	a ata	atgta	atta	ctad	cctgo	caa a	aaaa	aaaaa	973
)> 35															
	.> 86															
	2> D1 3> Ho	AA Omo s	sapie	ens												
<220) >															
	.> CI	os														
<222	2> 12	265	527													
		ig_pe 26]	_	de												
<223	3 > V	on He	eijne	e mai	crix											
		core eq II					3									
		olyA_		nal												
<222	2 > 8:	348	339													
		olyA_ 568		2												
)> 3!		etcai	tcate	ac to	cttto	gtago	e ata	ata	ette	tati	acto	ac a	aggag	caactt	60

actggaagaa ctcgtcatgc tctttgtagc gtggtgcttc tgttgctcac aggacaactt
gcctttgatg attttcaaga gagttgtgct atgatgtggc aaagtatgca ggaagcaggc
ggtca atg cct ctg gga gca agg atc ctt ttc cac ggt gtg ttc tat gcc
Met Pro Leu Gly Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala

								ctc Leu 5								218
								gar Glu								266
-	-	_	_	_				ctc Leu								314
	_		_					att Ile	_		_	_				362
								ggc Gly								410
-					_			cac His 85		-		_				458
								gtg Val								506
	_	-	gtg Val		_		taga	agaco	gac c	caga	agac	ec ca	agctt	gctt	:	557
ctag	gtcca	itc c	ettec	ctca	at ct	ctac	cata	a tgg	gccac	tgg	ggtg	gtgg	100 C	catct	cagtg	617
acag	gacac	ctc c	ctgca	acco	a gk	ttttc	cago	cac	cagt	ggg	atga	ıtggt	at g	gtgc	agcac	677
atgg	gtaat	tt t	ggtg	gtaat	t ct	aact	tggg	g cac	caaco	gaat	gcta	itttg	gtc a	tttt	taaac	737
tgaa	atccc	jaa a	igaaa	ctc	t at	tata	aatt	: taa	gata	atg	taat	gtat	tt g	gaaag	gtgctt	797
_	ataaa aaaaa		_	atga	it aa	aagg	gaato	aga	atta	ata	aaat	gttt	gt t	gato	tttaa	857 86 8

<210> 358 <211> 519 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 66..320 <221> sig_peptide <222> 66..113 <223> Von Heijne matrix score 3.5 seq TALAAXTWLGVWG/VR <221> polyA_signal <222> 490..495 <221> polyA_site <222> 508..519 <400> 358

aattagegeg taacgeasag actgettget geggeagaga egecagakgt geageteeag 60 cagca atg gca gtg acg gcg ttg gcg gcg mrg acg tgg ctt ggc gtg tgg Met Ala Val Thr Ala Leu Ala Ala Xaa Thr Trp Leu Gly Val Trp 110 -10 ggc gtg agg acc atg caa gcc cga ggc ttc ggc tcg gat cag tcc gag 158 Gly Val Arg Thr Met Gln Ala Arg Gly Phe Gly Ser Asp Gln Ser Glu 1

Asn Val Asp Arg Gly Ala Gly Ser Ile Arg Glu Ala Gly Gly Ala Phe 20 25 30	206
gga aag aga gag cag gct gaa gag gaa cga tat ttc cga gca cag agt Gly Lys Arg Glu Gln Ala Glu Glu Glu Arg Tyr Phe Arg Ala Gln Ser 35 40 45	254
aca gaa caa ctg gca rct ttg aaa aaa crc cat gaa gaa gar atc gtt Thr Glu Gln Leu Ala Xaa Leu Lys Lys Xaa His Glu Glu Glu Ile Val 50 55 60	302
cat cat aga gaa gga gat tgagcgtctg cagaaagaaa ttgagcgcca His His Arg Glu Gly Asp 65	350
taagcagaag atcaaaatgc tagaacatga tgattaagtg cacaccgtgt gccatagaat ggcacatgtc attgcccact tctgtgtaaa catggttctg gtttaactaa tatttgtctg tgtgctacta acagattata ataaattgtc atcagtgaaa aaaaaaaa	410 470 519
<210> 359 <211> 1028 <212> DNA <213> Homo sapiens	
<220>	
<221> CDS (1) (2) (2) (2) (2) (3) (4) (4) (4) (4) (4) (4) (4) (4) (4) (4	
<221> sig_peptide	
<pre><223> Von Heijne matrix score 4.40000009536743</pre>	
seq IVLHLVLQGMVYT/EY	
<221> polyA_site <222> 10161028	
<400> 359	
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagagag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac	60 111
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagagag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20	111
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagagag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20 cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5	111
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagaag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20 cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5 gag tac acc ttg gaa gta ttt ggc tac tgt cag gag ctg gag ttg tcc Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser	111
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagaag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20 cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5 gag tac acc ttg gaa gta ttt ggc tac tgt cag gag ctg gag ttg tcc Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser 1 5 10 15 ttg cat tac ctt ctt ctg ccc tat ctg ctg cta ggt gta aac ctg ttt	111
agetttaaag geetggeeag gggaggagea eagatatttt eetgtataat tecagaatgt etteagaag ee atg eat gga ttg ett eat tae ett tte eat aeg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20 cae ace tte att gte etg eac etg gte ttg eaa ggg atg gtt tat aet His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5 gag tae ace tgg gaa gta ttt gge tae tgt eag gag etg gag ttg tee Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser 1 5 10 15 ttg eat tae ett ett etg eec tat etg etg eta ggt gta aac etg ttt Leu His Tyr Leu Leu Leu Pro Tyr Leu Leu Leu Gly Val Asn Leu Phe 20 25 30	111 159 207 255
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagaag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20 cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5 gag tac acc tgg gaa gta ttt ggc tac tgt cag gag ctg gag ttg tcc Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser 1 5 10 15 ttg cat tac ctt ctt ctg ccc tat ctg ctg cta ggt gta aac ctg ttt Leu His Tyr Leu Leu Leu Pro Tyr Leu Leu Leu Gly Val Asn Leu Phe	111 159 207
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagagag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20 cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5 gag tac acc tgg gaa gta ttt ggc tac tgt cag gag ctg gag ttg tcc Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser 1 5 10 15 ttg cat tac ctt ctt ctg ccc tat ctg ctg cta ggt gta aac ctg ttt Leu His Tyr Leu Leu Leu Pro Tyr Leu Leu Leu Gly Val Asn Leu Phe 20 25 30 ttt ttc acc ctg act tgt gga acc aat cct ggc att ata aca aaa gca Phe Phe Thr Leu Thr Cys Gly Thr Asn Pro Gly Ile Ile Thr Lys Ala 35 40 45 aat gaa tta tta ttt ctt cat gtt tat gaa ttt gat gaa ktg atg ttt Asn Glu Leu Leu Phe Leu His Val Tyr Glu Phe Asp Glu Xaa Met Phe	111 159 207 255
agctttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt cttcagaag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn -25 -20 cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5 gag tac acc tgg gaa gta ttt ggc tac tgt cag gag ctg gag ttg tcc Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser 1 5 10 15 ttg cat tac ctt ctt ctg ccc tat ctg ctg cta ggt gta aac ctg ttt Leu His Tyr Leu Leu Leu Pro Tyr Leu Leu Leu Gly Val Asn Leu Phe 20 25 30 ttt ttc acc ctg act tgt gga acc aat cct ggc att ata aca aaa gca Phe Phe Thr Leu Thr Cys Gly Thr Asn Pro Gly Ile Ile Thr Lys Ala 35 40 45 aat gaa ttt tta ttt ctt cat gtt tat gaa ttt gat gaa ktg atg ttt	111 159 207 255 303

His	Cys	Val	Trp	Val	Asn	Asn	Cys	Ile 105	Gly	Ala	Trp	Asn	atc Ile 110	Arg	хаа	495
Phe	Leu	Ile 115	Tyr	Val	Leu	Thr	Leu 120	Thr	Ala	Ser	Ala	A1a 125	acc Thr	vai	Ala	543
Ile	Val	Ser	Thr	Thr	Phe	Leu 135	Val	His	Leu	Val	Val 140	Met	tca Ser	Asp	ьeu	591
tac Tyr 145	cag	gag Glu	act Thr	tac Tyr	atc Ile 150	gat Asp	gac Asp	ctt Leu	gga Gly	cac His 155	ctc Leu	cat His	gtt Val	atg Met	gac Asp 160	639
acq	gtc Val	ttt Phe	ctt Leu	att Ile 165	cag Gln	tac Tyr	ctg Leu	ttc Phe	ctg Leu 170	act Thr	ttt Phe	cca Pro	cgg Arg	att Ile 175	gtc Val	687
ttc Phe	atg Met	ctg Leu	ggc (Gly	ttt Phe	gtc Val	gtg Val	gtt Val	ctg Leu 185	arc Xaa	ttc Phe	ctc Leu	ctg Leu	ggt Gly 190	ggc	tac Tyr	735
ctg Leu	ttg Leu	ttt Phe 195	ata	cta	tat Tyr	ctg Leu	gcg Ala 200	gcc Ala	acc Thr	aac Asn	cag Gln	act Thr 205	act Thr	aac Asn	gag Glu	783
tgg Trp	tac Tyr 210	aga	rgt Xaa	gac Asp	tgg Trp	gcc Ala 215	tgg Trp	tgc Cys	cag Gln	cgt Arg	tgt Cys 220	ccc Pro	ctt Leu	gtg Val	gcc Ala	831
tgg Trp 225	cct	ccg Pro	tca Ser	gca Ala	gar Glu 230	Pro	caa Gln	gtc Val	cac His	cgg Arg 235	aac Asn	att I le	cac His	tcc Ser	cat His 240	879
aaa	ctt Leu	cgg Arg	arc Xaa	aac Asn 245	ctt	caa	gar Glu	atc Ile	ttt Phe 250	Leu	cct Pro	gcc Ala	ttt Phe	cca Pro 255	tgt Cys	927
				aaa Lys				cmag			ctgc	ct t	tgag	ctgt	a	978
gttcccgttt atttacacat gtggatcctc gttttccaaa aaaaaaaaaa																

<210> 360

<211> 452

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 69..434

<221> sig_peptide

<222> 69..236

<223> Von Heijne matrix
score 4.90000009536743
seq FACVPGASPTTLA/FP

<221> polyA_signal

<222> 419..424

<221> polyA_site

<222> 441..452

<400> 360

acagcgtgas tcgcccgcca gaagaatatg aaaaagcaga gcganctcgg ttaagggaaa 60 gcgccgag atg acg ggc ttt ctg ctg ccg ccc gca agc aga ggg act cgg 110 Met Thr Gly Phe Leu Leu Pro Pro Ala Ser Arg Gly Thr Arg

-55 -50 -4 5									
aga tca tgc Arg Ser Cys -40	agc aga	Ser Arg I	aaa agg Lys Arg -35	caa acg Gln Thr	aga aga Arg Arg -30	agg Arg	agg Arg	aac Asn	158
cca agt agc Pro Ser Ser -25	ttt gtg Phe Val	gct tcg (Ala Ser (tgt cca Cys Pro	acc ctc Thr Leu	ttg ccc Leu Pro -15	ttc (gcc Ala	tgt Cys	206
gtg cct gga Val Pro Gly	gcc agt Ala Ser	ccc acc a	acg ctc Thr Leu	gcg ttt Ala Phe 1	cct cct	vai.	ktg Xaa 5	ctc Leu	254
-10 aca ggt ccc Thr Gly Pro	avc acc Xaa Thr 10	gat ggc	att ccc Ile Pro 15	ttt gcc	ctr nak Leu Xaa	tct Ser . 20	gca Ala	gcg Ala	302
ggt ccc ttt Gly Pro Phe 25	tat act	Ser Phe	ccc tca	ggt avc Gly Xaa	ctc tct Leu Ser	ccc	cct Pro	gjà aaa	350
cca ctc ccg Pro Leu Pro	ggg gtg Gly Val	agg ggg	tta ccc	ctt ccc Leu Pro	agt gtt	ttt Phe	tat Tyr	tcc Ser	398
40 tgt ggg gct Cys Gly Ala	cac ccc His Pro	aaa gta Lys Val	tta aaa Leu Lys	gta gct Val Ala 65	ttg taa	ttcaa	aa		444
55 a aa aaaaa		60		03					452
<210> 361									
<211> 875 <212> DNA <213> Homo	sapiens					•			
<220> <221> CDS	•								
<222> CDS <222> 628	804								
<221> sig_p <222> 628 <223> Von H	711	trix							
score	4.19999 MPVIPALQI	980926514	ŧ	·					
<221> polyA <222> 864	_								
<400> 361 aaagatggac	accgcgga	gg aagaca	atatg tag	gagtgtgt	cggtcag	aag g	gaaca	acctga	60
gaaaccgctt	tatcatcc	tt gtgtat	gtac tg	gcagtatt	aagttng	itcc a	atcaa	agaatg	120 180
cttagttcaa tgcttttaca	ccaattta	ac acagto tt ctccao	gaaa ag gatat qc	aatactgt cttcacqq	cttccaa	ttc a	agcad	catatt	240
tqctqqactq	gttacaag	ta ttggca	actgc aa	tacgatat	tggtttc	att a	ataca	acttgt	300 360
ggcctttgca ggagttattt	tggttggg	ag ttgtto	ctct ta	cagcatgt	gagtatt	cat q	gcct	ctgatt	420
tatttgactg	atgtttag	tt atttga	atgtc ag	agtgtcat	gtattag	ggaa a	agcci	ttactt	480
araaratqtt	catcggaa	ct aaraat	tgakt tt	aacaggto	: agttttt	tga 🤉	gtgaa	atgtgg	540 600
gaaaraacac aaatcaaatc	agcataca ataattag	ga atgget at atgaag	gt atg c	ta rag c eu Xaa L	tt tca a leu Ser A	agg go	ct a	ca aaa	654
rac ggc cgc Xaa Gly Arc	g Ala Arg	Trp Leu	atg cct Met Pro	gta ato	cca gca	a ctt a Leu	cag Gln -5	gag	702
gcc gan gca	-15 ggc gga		ggt cag		gaa act	agc		gcc	750

Ala Xaa Ala Gly Gly Ser Arg Gly Gln Glu Phe Glu Thr Ser Leu Ala	
aac atg gag act gag gca gga gaa ttg ctt aaa ccc agg agg cgg agg Asn Met Glu Thr Glu Ala Gly Glu Leu Leu Lys Pro Arg Arg Arg Arg 15 20 25	798
ttg car tgaactgaga tcgcaccact gcactccage ttgggcaaca gagcaagact Leu Gln	854
30 ttgtctcgca aaaaaaaaa a	875
<210> 362 <211> 531 <212> DNA	
<213> Homo sapiens	
<220>	
<221> CDS <222> 70366	
<221> sig_peptide <222> 70108	
<223> Von Heijne matrix	
score 3.5 seq MHLLSNWANPASS/RR	
<221> polyA_signal	
<221> polyA_site	
<222> 521531	
<400> 362 aagtggccat ggcggataca gcgactacag catcggcggc ggcggctagt gccgctag	gcg 60
cctcgagcg atg cac ctc ctt tcc aac tgg gca aac ccc gct tcc age of Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser	aga +++
cot cot tot ato goo got toa ggo act tot tgg ata toa tog acc ot	c 159
Arg Pro Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Le	1
gca cac tot ttg toa ctg aga gac gto toa gag agg ctg tgc ago tg Ala His Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cy	c 207 s
20 25 30 tog agg act ata agg atg gga ccc tgc gcc cgg ggg tca cca atg aa	c 255
Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met As	11
ago tot gga gtg cac aga aaa toa ago agg ota tto tac ato ogg ac Ser Ser Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Th	a 303 r
50 55 60 63	
cca atg aga aga tct tca tgc cat tta gaa tgt crg gtt ata ttc ct Pro Met Arg Arg Ser Ser Cys His Leu Glu Cys Xaa Val Ile Phe Le	u
ttg gga cgc caa ttg taaktgttac cttcaaagga tttccttttc taaaaaatt	a 406
Leu Gly Arg Gln Leu 85	
ttttaratgt ctaactttat gttattgctc acgggtattt gactgaattg ttgattt ataagtcaat tcctggaggg aaattaccaa ataaaatgat atgtatttct taccaca	agg 466 aaa 526
aaaaa	531

```
<210> 363
<211> 1244
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 70..366
<221> sig_peptide
<222> 70..108
<223> Von Heijne matrix
     score 3.5
     seq MHLLSNWANPASS/RR
<221> polyA site
<222> 1233..1244
<400> 363
aagtggccat ggcggataca gcgactacag catcggcggc ggcggctagt gccgctagcg
cctcgagcg atg cac ctc ctt tcc aac tgg gca aac ccc gct tcc agc aga
          Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg
                      -10
                                          -5
                                                                      159
cgt cct tct atg gcc gct tca ggc act tct tgg ata tca tcg acc ctc
Arg Pro Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu
                                                                      207
qca cac tot ttq toa ctq aqa qac qtc tca gag agg ctg tgc agc tgc
Ala His Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys
                            25
                                                30
tgg agg act ata agc atg gga ccc tgc gcc cgg ggg tca cca atg aac
                                                                      255
Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn
   35
                        40
                                            45
                                                                      303
age tet gga gtg cae aga aaa tea age agg eta tte tae ate egg aca
Ser Ser Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr
                    55
                                        60
cca atg aga aga tot toa tgc cat tta raa tgt cag gtt ata ttc ctt
                                                                      351
Pro Met Arg Arg Ser Ser Cys His Leu Xaa Cys Gln Val Ile Phe Leu
ttg gga cgc caa ttg tagtcggtct tctcttgccc aaccagacac tggcatccac
                                                                      406
Leu Gly Arg Gln Leu
tgtcttctgg cagtggctga accagagcca caatgcctgt gtcaactatg caaaccgcaa
                                                                     466
                                                                     526
tgcraccaag ccttcacctg catccaagtt catccaggga tacctgggag ctgtcatcag
                                                                     586
cgccgtctcc attgctgtgg gccttatktc ctggttcaga aagccaacaa gttcaccca
                                                                      646
gccaccegcc ttctcatcca gaggtttgtg ccgttccctg ctgtagccag tgccaatatc
                                                                     706
tgcaatgtgg tcctgatgcg gtacggggag ctggaggaag ggattgatgt cctggacagc
                                                                     766
gatggcaacc tcgtgggctc ctccaagatc gcagcccgac acgccctgct ggagacggcg
                                                                     826
ctgacgcgag tggtcctgcc catgcccatc ctggtgctac ccccgatcgt catgtccatg
                                                                     886
ctggagaaga cggctctcct gcaggcacgc ccccggctgc tcctccctgt gcaaagcctc
                                                                     946
gtgtgcctgg cagcettegg cetggecetg cegetggeca teageetett ceegcaaatg
tcagagattg aaacatccca attagagccg gagatagccc aggccacgag cagccggaca
                                                                    1006
gtggtgtaca acaaggggtt gtgagtgtgg tcagcggcct ggggacggag cactgtgcag
                                                                    1066
                                                                    1126
ccggggagct gaggggcarg gccgtagact cacggctgca cctgcaggga gcagcacgcc
aaccccagca gtcctgggcc ccctgggaga gtgctcaacc tacagtggag ggagactgac
                                                                    1186
                                                                     1244
ccattcacat tttaacatag gcaagaggag ttctaacaca tttcgtacaa aaaaaaaa
```

<210> 364

<211> 631

<212> DNA

<213> Homo sapiens

```
<220>
<221> CDS
<222> 111..434
<221> sig_peptide
<222> 111..185
<223> Von Heijne matrix
      score 3.90000009536743
      seq WIAAVTIAAGTAA/IG
<221> polyA site
<222> 618..631
<400> 364
aatogoggag toggtgottt agtacgoogo tggcacettt actotogoog googoggaa
                                                                      60
cccgtttgag ctcggtatcc tagtgcacac gccttgcaag cgacggcgcc atg agt
                                                                     116
                                                       Met Ser
ctg act tcc agt tcc agc gta cga gtt gaa tgg atc gca gca gtt acc
                                                                     164
Leu Thr Ser Ser Ser Ser Val Arg Val Glu Trp Ile Ala Ala Val Thr
            -20
att gct gct ggg aca gct gca att ggt tat cta gct tac aaa aga ttt
                                                                     212
Ile Ala Ala Gly Thr Ala Ala Ile Gly Tyr Leu Ala Tyr Lys Arg Phe
tat gtt aaa gat cat cga aat aaa gct atg ata aac ctt cac atc cag
                                                                     260
Tyr Val Lys Asp His Arg Asn Lys Ala Met Ile Asn Leu His Ile Gln
                   15
                                       2.0
aaa gac aac ccc aag ata gta cat gct ttt gac atg gag gat ttg gga
                                                                     308
Lys Asp Asn Pro Lys Ile Val His Ala Phe Asp Met Glu Asp Leu Gly
               30
                                   35
gat aaa get gtg tae tge egt tgt tgg agg tee aaa aag tte eea tte
                                                                     356
Asp Lys Ala Val Tyr Cys Arg Cys Trp Arg Ser Lys Lys Phe Pro Phe
                                50
           4.5
tgt gat ggg gct cac aca aaa cat aac gaa gag act gga gac aat gtg
                                                                     404
Cys Asp Gly Ala His Thr Lys His Asn Glu Glu Thr Gly Asp Asn Val
       60
                          65
                                               70
ggc cct ctg atc atc aag aaa aaa gaa act taaatggaca cttttgatgc
                                                                     454
Gly Pro Leu Ile Ile Lys Lys Lys Glu Thr
                       80
tgcaaatcag cttgtcgtga agttacctga ttgtttaatt araatgacta ccacctctgt
                                                                     514
ctgattcacc ttcgctggat tctaaatgtg gtatattgcm aactgcagct ttcacattta
                                                                     574
tggcatttgt cttgttgaaa catcgtggtg cacatttgtt taaacaaaaa aaaaaaa
```

<210> 365

<211> 781

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 19..567

<221> sig_peptide

<222> 19..63

<223> Von Heijne matrix
score 8.39999961853027
seq AMWLLCVALAVLA/WG

<221> polyA_signal

<222> 749..754

<221> polyA_site <222> 771..781

<400> 365 51 aagtgctgct tacccatc atg gaa gca atg tgg ctc ctg tgt gtg gcg ttg Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu -10 gcg gtc ttg gca tgg ggc ttc ctc tgg gtt tgg gac tcc tca gaa cga Ala Val Leu Ala Trp Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg 10 atg aag agt cgg gag cag gga aga cgg ctg gga gcc gaa agc cgg acc 147 Met Lys Ser Arg Glu Gln Gly Arg Arg Leu Gly Ala Glu Ser Arg Thr 20 25 195 ctg ctg gtc ata gcg cac cct gac gat gaa gcc atg ttt ttt gct ccc Leu Leu Val Ile Ala His Pro Asp Glu Ala Met Phe Phe Ala Pro aca gtg cta ggc ttg gcc cgc cta agg cac tgg gtg tac ctg ctt tgc 243 Thr Val Leu Gly Leu Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys 55 50 ttc tct gca gga aat tac tac aat caa gga gag act cgt aag aaa gaa 291 Phe Ser Ala Gly Asn Tyr Tyr Asn Gln Gly Glu Thr Arg Lys Lys Glu 65 70 ctt ttg car age tgt gat gtt ttg ggg att cca ctc tcc agt gta atg 339 Leu Leu Gln Ser Cys Asp Val Leu Gly Ile Pro Leu Ser Ser Val Met 85 387 att att gac aac agg gat ttc cca rat gac cca ggc atg cag tgg gac Ile Ile Asp Asn Arg Asp Phe Pro Xaa Asp Pro Gly Met Gln Trp Asp 95 100 aca rag cac gtg gcc ara gtc ctc ctt cag cac ata gaa gtg aat ggc 435 Thr Xaa His Val Ala Xaa Val Leu Leu Gln His Ile Glu Val Asn Gly 115 120 483 atc aat ctg gtg gtg act ttc gat gca ggg gga rta agt ggc cac agc Ile Asn Leu Val Val Thr Phe Asp Ala Gly Gly Xaa Ser Gly His Ser 135 125 130 aat cac att gct ctg tat gca gct gtg agg aag ctt gag ggc caa att 531 Asn His Ile Ala Leu Tyr Ala Ala Val Arg Lys Leu Glu Gly Gln Ile 145 150 tgc aag ccc tgt ggc act gga caa gac ttt aag gaa tgagtgctgt 577 Cys Lys Pro Cys Gly Thr Gly Gln Asp Phe Lys Glu 160 caatcagtgt gcctccacct tcaccatctt cttcccctta ctctcacttc cgtcatgtgt 637 697 tttatacaac tctcaaatct ttcttqqaqa aqqagqatat acatacataa tatgaaatgt 757 gtttgttctt cacagtcacc cgattttact gatatttatt tgcattttac caataaaaag

781

<210> 366

<211> 931

<212> DNA

<213> Homo sapiens

aaaatgcaag ctcaaaaaaa aaaa

<220>

<221> CDS

<222> 19..312

<221> sig_peptide

<222> 19..63

PCT/IB98/02122

seq AMWLLCVALAVLA/WG

<221> polyA signal <222> 896..901 <221> polyA site <222> 921..931 <400> 366 aagtgctgct tacccatc atg gaa gca atg tgg ctc ctg tgt gtg gcg ttg Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu

99 geg gtc ttg gca tgg ggc ttc ctc tgg gtt tgg gac tcc tca gaa cga Ala Val Leu Ala Trp Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg atg aag agt cgg gag cag gga rga cgg ctg gga gcc gaa agc cgg acc 147 Met Lys Ser Arg Glu Gln Gly Xaa Arg Leu Gly Ala Glu Ser Arg Thr 15 ctg ctg gtc ata gcg cac cct gac gat gaa gcc atg ttt ttt gct ccc 195

-10

Leu Leu Val Ile Ala His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro 243 aca gtg cta ggc ttg gcc cgc cta agg cac tgg gtg tac ctg ctt tgc Thr Val Leu Gly Leu Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys 50 55 60 291

ttc tct gca gtt ttc cgt agg gag cta agt gaa tac acc gaa rgt ctt Phe Ser Ala Val Phe Arg Arg Glu Leu Ser Glu Tyr Thr Glu Xaa Leu 70 65 342

acc tot gaa coc ctc ama goc tagggacagg arcggccggc ttacctggtg Thr Ser Glu Pro Leu Xaa Ala

ggttggggga cgtcggcagc tcrcgtacta cgccagcagg attganganc acagaaacag 402 462 ttgchsttgg ttgtattcag tacctkcatt tccgttggga actccaccwg tacttgttat 522 kctgtggaac ttttttttat ttgtagaagg agcaagaata ttgaccttac tatatagcac 582 acgaaacaat ctatgctgta tcgtgcctgc tcaatcctta aagttaactt ctaatgatag 642 taaaaracct tootgotgoo tttaaaaatgo agottgtgot aktaacatgo atgtgtcaaa 702 ttgaaraatt agacatagat gactaratar aaagtaattt tgtaggtaat tttaragttc 762 aactccaccc agctttcakt gaaggaacct ttcaaataat aratttttgc ttaccatara raaaaratca aatgacaaag caaatattga ccattaagct ggaatatggt gataattgaa 822

cagttgtata aatgaaktaa ttgaattgta cacatacaat gggtgaattt tatggcatgt 882 caaagtatac ctcaataaag ctatttttt aaattgcmaa aaaaaaaaa

<210> 367

<211> 849

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 64..612

<221> sig_peptide

<222> 64..234

<223> Von Heijne matrix score 3.79999995231628 seq QLWLVMEFCGAGS/VT

<221> polyA site

<222> 839..849

<400> 367

acatacgggc aagtttataa gggtcgtcat gtcaaaacgg gccagcttgc agccatcaag gtt atg gat gtc aca ggg gat gaa gaa gaa atc aaa caa gaa att														60		
att	ata	cat	atc	aca	aaa	gat	gaa	qaq	qaa	gaa	atc	aaa	Caa	gaa	acc	108
900	Met	Asp	Val	Thr	Gly	Asp	Glu	Glu	Glu	Glu	Ile	Lys	Gln	Glu	·Ile	
			-55					-50					-45			
aac	atq	ttg	aag	aaa	tat	tct	cat	cac	cgg	aat	att	gct	aca	tac	tat	156
Asn	Met	Leu	Lys	Lys	Tyr	Ser	His	His	Arg	Asn	Ile	Ala	Thr	Tyr	Tyr	
		-40					-35					- 30				204
ggt	gct	ttt	atc	aaa	aag	aac	cca	cca	ggc	atg	gat	gac	caa	CCC	tgg	204
Gly	Ala	Phe	Ile	Lys	Lys	Asn	Pro	Pro	Gly	Met	Asp	Asp	Gln	Leu	Trp	
	-25					-20					-15					252
ttg	gtg	atg	gag	ttt	tgt	ggt	gct	ggc	tct	gtc	acc	gac	ctg	acc	aay	232
Leu	Val	Met	Glu	Phe		Gly	Ala	Gly	Ser	Val	Thr	Asp	Leu	5	цуз	
-10					- 5					1		~	+ > 0	-	tac	300
aac	aca	aaa	ggt	aac	acg	ttg	aaa	gag	gag	tgg	att	yca Nla	tac	Tle	Cve	300
Asn	Thr	Lys		Asn	Thr	Leu	гЛs	GIU	GIU	Trp	116	Ala	Tyr 20	110	Cyb	
			10					15		a o a	~ ~ ~	cat		ata	att	348
msg	gaa	atc	tta	cgg	999	ctg	art	cac	Tou	Tic	Gln	His	aaa	Val	Ile	
Xaa	Glu		Leu	Arg	GIY	ьeu	лаа	urs	neu	nis	Gii	35	Lys			
		25					30	a+ c	++-	cta	act		aat	aca	gaa	396
cat	cga	rat	att	aaa	999	Caa	Aac	77-1	T.em	Leu	Thr	Glu	aat Asn	Ala	Glu	
His		хаа	Tie	гÀг	GIY	45	ASII	val	ne u	шец	50	914	11011			
	40		~~	~~~			rt.c	ak+	act	cad		gat	cga	aca	ata	444
gtt	aaa	cta	grg	gac	Dho	Glv	Yaa	Yaa	Δla	Gln	Leu	Asp	Arg	Thr	Val	
	пуѕ	Dear	val	ASP	60	Giy	nau	nuu	1120	65					70	
55	200	220	+	a c t		att	gga	act	ccc		taa	atq	gca	cca	raa	492
990	. ayy	Xaa	Asn	Thr	Phe	Ile	Glv	Thr	Pro	Tyr	Trp	Met	Ala	Pro	Xaa	
Gry	7.19	nuu		75			2		80	•	_			85		
att	att	acc	tat	gat	gaa	aac	сса	sat	gcc	aca	tat	gat	tto	aar	art	540
Val	Ile	Ala	Cvs	Asp	Glu	Asn	Pro	Xaa	Āla	Thr	Tyr	Asp	Phe	Lys	Xaa	
			90					95					100			
gac	ttq	tqq	tct	ttg	ggt	ato	acc	gco	att	gaa	. atg	gca	. gaa	999	ctc	588
Asp	Leu	Trp	Ser	Leu	Gly	Ile	Thr	Ala	. Ile	Glu	Met	. Ala	. GIU	. Gly	Leu	
		105					110					115				640
ccc	cto	tct	gtg	aca	tgo	acc	сса	. tga	gago	tct	cttc	ctca	tc c	cccg	gaatc	642
Pro	Leu	Ser	· Val	Thr	Cys	Thr	Pro)								
	120					125										702
cag	gcgcc	tcg	gctg	aagt	ct a	agaa	gtgg	rt ca	aaaa	aatt	. cca	igtca	ובננ	attg	agagct	
act	taat	aaa	aaat	caca	ac c	agco	racca	q ca	lacas	gaaca	att	gate	aag	catc	callla	
tac	gaga	cca	acct	aatg	ag c	gaca	iggto	c go	atto	aact	caa	iggac	cat	acco	atagaa	849
caa	agaa	igaa	gcga	ıggaa	aa a	ıaaaa	ıaa									

<210> 368 <211> 644

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 39..458

<221> sig_peptide <222> 39..80

<223> Von Heijne matrix score 4.40000009536743 seq FLTALLWRGRIPG/RQ

<221> polyA_signal <222> 613..618

<221> polyA_site <222> 633..644

<pre><400> 368 agcggagacg cagagtcttg agcagcgcgn caggcacc atg ttc ctg act gcg ctc</pre>														56	
ctc tg Leu Tr													cac	cgg	104
cgg cc Arg Pr 10	g cgg o Arg	ttc Phe	gtg Val	tcg Ser	ttg Leu 15	cgc Arg	gcc Ala	aag Lys	cag Gln	aac Asn 20	atg Met	atc Ile	cgc Arg	cgc Arg	152
ctg ga Leu Gl 25	g atc u Ile	gag Glu	gcg Ala	gag Glu 30	aac Asn	cat His	tac Tyr	tgg Trp	ctg Leu 35	agc Ser	atg Met	ccc Pro	tac Tyr	atg Met 40	200
acc cg Thr Ar	g gag g Glu	cag Gln	gag Glu 45	cgc Arg	ggc Gly	cac His	gcc Ala	gcg Ala 50	ttg Leu	cgc Arg	agg Arg	agg Arg	gag Glu 55	gcc Ala	248
ttc ga Phe Gl	u Ala	Ile 60	Lys	Ala	Ala	Ala	Thr 65	Ser	Lys	Phe	Pro	Pro 70	His	Arg	296
ttc at Phe Il	t gcg e Ala 75	gac Asp	cag Gln	ctc Leu	gac Asp	cat His 80	ctc Leu	aat Asn	vgt Xaa	cac His	caa Gln 85	gaa Glu	atg Met	gtc Val	344
cta at Leu Il 90	e Leu	agt Ser	cgt Arg	cac His	cct Pro 95	tgg Trp	att Ile	tta Leu	tgg Trp	atc Ile 100	acg Thr	gag Glu	ctg Leu	acc Thr	392
atc tt Ile Ph 105	t acc e Thr	tgg Trp	tct Ser	gga Gly 110	ctg Leu	aaa Lys	aac Asn	tgt Cys	agc Ser 115	ttg Leu	tgt Cys	gaa Glu	aat Asn	gag Glu 120	440
ctt tg Leu Tr					taaa	aacaa	aac a	aaaca	atgag	gt ag	gtct	gcata	a		488
tcgaat tcatta ctgcaa	gtct	gata	ggaa	ga ta	aggga	attt	c cto	cagto	aagt caca	ctaa	attgo gata	etg 1	tcctq	gtggtt ggaaag	548 608 644

<210> 369

<211> 918

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 9..185

<221> sig_peptide

<222> 9..50

<223> Von Heijne matrix
score 3.70000004768372
seq AALVTVLFTGVRR/LH

<221> polyA_site <222> 906..918

<400> 369
agctcagc atg gct gct tta gtg act gtt ctc ttc aca ggt gtc cgg agg
Met Ala Ala Leu Val Thr Val Leu Phe Thr Gly Val Arg Arg
50

- 5

-10

ctg cac tgc agc gcr scg ctt ggg cgg gcg gcc agt ggc grc tac a Leu His Cys Ser Ala Xaa Leu Gly Arg Ala Ala Ser Gly Xaa Tyr S	
agg aac tgg ctg cca acc cct ccg gct acg ggc ccc tta ccg agc t Arg Asn Trp Leu Pro Thr Pro Pro Ala Thr Gly Pro Leu Pro Ser S 20 25 30	
cag act ggt cat atg cgg atg gcc gcc ctg ctc ccc caa tgaaaggcc Gln Thr Gly His Met Arg Met Ala Ala Leu Leu Pro Gln 35 40 45	a 195
gcttcgaaaa aaagctgaaa gggagacktt tgcaaracra kttgtactgc tgtcac	agga 255
aatggacgct ggattacaas catggcasct caggcagcar aakttgcagg aaraac	
gaagcaggaa aatgctctta aacccaaagg ggcttcactg aaaascccac ttccaa	
ataaaaagca actectgeet ecetteetea ecetgtetet ggatttettt tetate	
aratgettea tecagecara aaatageett cackkteece atetgtette aragea	
agotgggacm ccaaraacaa gotgttarat cactgootgg gaggottggo ttarta	
catctotggt tocattocag ttoagotaag tottgottta aaatttttac otoota	
ggtgcggtgg ctcacgcctg taatcccagc actttgggag gctgaggcgg gcagat	
 agatcaggag ttcgagacca gcctggccaa cccagcctgg tcaacatggt gaaacc ccctactaaa gatacaaaca attagccggg cgtggtgggg tgcgcttgta atccca 	-
	J
ctcaggaggc tgaggcagga gaatcgctta aactcgggag gtagaggttg cagtga	_
aggtcacacc attgcactcc aacctgggcg acagggcgag actctgtctc aaaaaa	
aaa	918
<pre><210> 370 <211> 472 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 14316 <221> sig_peptide <222> 14121 <223> Von Heijne matrix</pre>	
attatataga gcc atg ggg cct tac aac gtg gca gtg cct tca gat gta Met Gly Pro Tyr Asn Val Ala Val Pro Ser Asp Val -35 -30	l 5
tct cat gcc cgc ttt tat ttc tta ttt cat cga cca tta agg ctg tt Ser His Ala Arg Phe Tyr Phe Leu Phe His Arg Pro Leu Arg Leu Le -20 -15 -10	
aat ctg ctc atc ctt att gag ggc agt gtc gtc ttc tat cag ctc ta Asn Leu Leu Ile Leu Ile Glu Gly Ser Val Val Phe Tyr Gln Leu Ty -5 1 5	
tcc ttg ctg cgg tcg gag aag tgg aac cac aca ctt tcc atg gct ct Ser Leu Leu Arg Ser Glu Lys Trp Asn His Thr Leu Ser Met Ala Le 10 15 20	
atc ctc ttc tgc aac tac tat gtt tta ttt aaa ctt ctc cgg gac ag	ga 241
Ile Leu Phe Cys Asn Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Ar 25 30 35 40	g

• •														
wta kta tta ggc agg gca tac tcc tac cca ctc aac agt tat gaa ctc Xaa Xaa Leu Gly Arg Ala Tyr Ser Tyr Pro Leu Asn Ser Tyr Glu Leu 45 50 55	289													
aag gca aac twa gct gcc tct caw caa tgagggagaa ctcagataaa Lys Ala Asn Xaa Ala Ala Ser Xaa Gln 60 65	336													
aatattttca tacgttctat ttttttcttg tgatttttat aaatatttaa gatattttat attttgtata ctattatgtt ttgaaagtcg ggaagagtaa gggatattaa atgtatccgt aaacaaaaaa aaaaam														
<210> 371 <211> 1504 <212> DNA <213> Homo sapiens														
<220> <221> CDS <222> 701092														
<pre><221> sig_peptide <222> 70234 <223> Von Heijne matrix</pre>														
<pre><221> polyA_signal <222> 14751480</pre>														
<221> polyA_site <222> 14931504														
<400> 371 agaaatcgta ggacttccga aagcagcggc ggcgtttgct tcactgcttg gaagtgtgag tgcgcgaag atg cga aag gtg gtt ttr att acc ggg gct agc agt ggc att Met Arg Lys Val Val Leu Ile Thr Gly Ala Ser Ser Gly Ile														
ggc ctg gcc ctc tgc aag cgg ctg ctg gcg gaa gat gat gag ctt cat Gly Leu Ala Leu Cys Lys Arg Leu Leu Ala Glu Asp Asp Glu Leu His -40 -35 -30	159													
ctg tgt ttg gcg tgc agg aat atg agc aag gca gaa gct gtc tgt gct Leu Cys Leu Ala Cys Arg Asn Met Ser Lys Ala Glu Ala Val Cys Ala -25 -20 -15 -10	207													
gct ctg ctg gcc tct cac ccc act gct gag gtc acc att gtc cag gtg Ala Leu Leu Ala Ser His Pro Thr Ala Glu Val Thr Ile Val Gln Val -5 1 5	255													
gat gtc agc aac ctg cag tca ttc ttc cgg gcc tcc aag gaa ctt aag Asp Val Ser Asn Leu Gln Ser Phe Phe Arg Ala Ser Lys Glu Leu Lys 10 15 20	303													
caa agg ttt cag aga tta gac tgt ata tat cta aat gct ggg atc atg Gln Arg Phe Gln Arg Leu Asp Cys Ile Tyr Leu Asn Ala Gly Ile Met 25 30 35	351													
cct aat cca caa cta aat atc aaa gca ctt ttc ttt ggc ctc ttt tca Pro Asn Pro Gln Leu Asn Ile Lys Ala Leu Phe Phe Gly Leu Phe Ser 40 45 50 55	399													
aga aaa gtg att cat atg ttc tcc aca gct gaa ggc ctg ctg acc cag Arg Lys Val Ile His Met Phe Ser Thr Ala Glu Gly Leu Leu Thr Gln 60 65 70	447													
ggt gat aag atc act gct gat gga ctt cag gag gtg ttt gag acc aat Gly Asp Lys Ile Thr Ala Asp Gly Leu Gln Glu Val Phe Glu Thr Asn 75 80 85	495													

gtc Val	ttt Phe	ggc Gly 90	cat His	ttt Phe	atc Ile	ctg Leu	att Ile 95	cgg Arg	gaa Glu	ctg Leu	gag Glu	cct Pro 100	ctc Leu	ctc Leu	tgt Cys	543
											tca Ser 115					591
											cac His					639
Glu	Pro	Tyr	Ser	Ser 140	Ser	Lys	Tyr	Ala	Thr 145	Asp		Leu	Ser	Val 150	Ala	687
ttg Leu	aac Asn	agg Arg	aac Asn 155	ttc Phe	aac Asn	cag Gln	cag Gln	ggt Gly 160	ctc Leu	tat Tyr	tcc Ser	aat Asn	gtg Val 165	gcc Ala	tgt Cys	735
cca Pro	ggt Gly	aca Thr 170	gca	ttg Leu	acc Thr	aat Asn	ttg Leu 175	aca Thr	tat Tyr	gga Gly	att Ile	ctg Leu 180	cct Pro	ccg Pro	ttt Phe	783
ata Ile	tgg Trp 185	acg	ctg Leu	ttg Leu	atg Met	ccg Pro 190	gca Ala	ata Ile	ttg Leu	cta Leu	ctt Leu 195	cgc Arg	ttt Phe	ttt Phe	gca Ala	831
	gca					cca					gaa Glu					879
ctt					cct						ctg Leu					927
agt Ser	gcc Ala	acc Thr	act Thr 235	ggc	ttt Phe	gga Gly	aga Arg	aat Asn 240	tac	att Ile	atg Met	acc Thr	cag Gln 245	aag Lys	atg Met	975
			gaa					aaa			caa Gln					1023
ctg Leu	gaa Glu 265	aag	cac His	att Ile	agg Arg	gtc Val 270	act Thr	att Ile	caa Gln	aaa Lys	aca Thr 275	gat Asp	aat Asn	cag Gln	gcc Ala	1071
Arg	ctc	_			Cys	cta	taai	ttcca	agc a	actt	ggga	ag go	ccaa	ggcag	B	1122
aagg ctac ctc aga tat tgt	Arg Leu Ser Gly Ser Cys Leu 280 285 aaggatcact tgagaccagg agttcaagac cagcctgaga aacatagtga gcccttgtct ctacaaaaag aaataaaaat aatagctggg tgtggtggca tgcgcatgta gtcccagcta ctcagaagga tgaggtggga ggatctcttg aggctgggag gcagaggttg cagtgagctg agattgtgcc actgcactcc agcctgggtg acagcgagac cctgtctcaa aatatgtata tatttaatat atatataaaa ccagagctga caatgacact ctggaacatt gcataccttc tgtacattct ggggtacatg gatttctact gagttggata atatgcattt gtaataaact atgaactatg aaaaaaaaaa aa														a 1242 g 1302 a 1362 c 1422	

. ____

<210> 372

<211> 765

<212> DNA

<213> Homo sapiens

<220>

.....

<221> CDS

<222> 274..597

<221> sig_peptide

<222> 274..399

<223> Von Heijne matrix score 5.19999980926514

seq LLFDLVCHEFCQS/DD

<221> polyA signal <222> 731..736 <221> polyA site <222> 754..765 <400> 372 accaggaaca tccagctatt tatgatagca tttgcttcat tatgtcaagt tcaacaaatg ttgacttgct ggtgaaggtg ggggaggttg tggacaagct ctttgatttg gatgagaaac 120 taatgttaag aatgggtcag aaatggggct gctcagcctc tggaccaacc ccaggaagag 180 totgaagago agcoagtgtt toggottgtg cootgtatac ttgaagotgo caaacaagta 240 cgttctgaaa atccagaatg gcttgatgtt tac atg cac att tta caa ctg ctt 294 Met His Ile Leu Gln Leu Leu act aca gtg gat gat gga att caa gca att gta cat tgt cct gac act 342 Thr Thr Val Asp Asp Gly Ile Gln Ala Ile Val His Cys Pro Asp Thr -25 -30 gga aaa gac att tgg aat tta ctt ttt gac ctg gtc tgc cat gaa ttc 390 Gly Lys Asp Ile Trp Asn Leu Leu Phe Asp Leu Val Cys His Glu Phe -10 -15 tgc cag tct gat gat cca gcc atc att ctt caa raa car aaa acr gtg 438 Cys Gln Ser Asp Asp Pro Ala Ile Ile Leu Gln Xaa Gln Lys Thr Val cta gcc tct gtt ttt tca gtg ttg tct gcc atc tat gcc tca cag act 486 Leu Ala Ser Val Phe Ser Val Leu Ser Ala Ile Tyr Ala Ser Gln Thr 25 534 gag caa gak tat cta aar ata raa aaa gga gac ggt ggc tca ggg agt Glu Gln Xaa Tyr Leu Lys Ile Xaa Lys Gly Asp Gly Gly Ser Gly Ser 35 40 aaa gga agg cca ktt gan caa aca gaa ktg ttc ctc tgc att tca aaa Lys Gly Arg Pro Xaa Xaa Gln Thr Glu Xaa Phe Leu Cys Ile Ser Lys 50 55 637 cot tot toc ttt cta tagcoctgtg gtggaagatt ttattaaaat cotacgtgaa Pro Ser Ser Phe Leu gttgataagg cgcttgctga tgacttggaa aaaaacttcc caagtttgaa ggttcagact 697 757 taaaacctga attggaatta cttctgtaca agaaataaac tttatttttc tcactgacaa 765 aaaaaaaa

<210> 373

<211> 1041

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 230..469

<221> sig_peptide

<222> 230..307

<223> Von Heijne matrix
 score 4.90000009536743
 seq VLCTNQVLITARA/VP

<221> polyA_signal

<222> 1004..1009

<221> polyA_site

<222> 1027..1040

<pre><400> 373 aacttccaag ttgtagtgtt gttgttttca gcctgctgct gctgctgcta ttgcggctag gggaaccgtc gtggggaagg atggtgtgcg aaaaatgtga aaagaaactt ggtactgtta</pre>														
tcactccaga tacatggaaa gatggtgcta ggaataccac agaaagtggt ggaagaaag	a 120 c 180													
tgaatgaaaa taaagctttg acttcaaaaa aagccagaat tgatccata atg gaa ga	a 238													
met Giu Giu -25														
ata agt tot coa ott gta gaa ttt gta aaa gtt ttg tgc acc aac cag	286													
Ile Ser Ser Pro Leu Val Glu Phe Val Lys Val Leu Cys Thr Asn Gln														
-20 -15 -10														
gtt ctc att act gcc agg gct gtg cct aca aaa aag gca tct gtg cga	334													
Val Leu Ile Thr Ala Arg Ala Val Pro Thr Lys Lys Ala Ser Val Arg														
-5 1 5	382													
tgt gtg gaa aaa agg ttt tgg ata cca aaa act aca agc aaa cat ctg	302													
Cys Val Glu Lys Arg Phe Trp Ile Pro Lys Thr Thr Ser Lys His Leu														
10 15 20	430													
tot aga tgt att gat gga att tot ggo ttt ota aat gat ttt act tto	150													
Ser Arg Cys Ile Asp Gly Ile Ser Gly Phe Leu Asn Asp Phe Thr Phe														
30 35 40 tgc ctt gaa ttt tca agg cat aga tgt caa ctt aca gaa taacatgtkt	479													
Cys Leu Glu Phe Ser Arg His Arg Cys Gln Leu Thr Glu														
45 50														
taagataatt aagtktaaac cagaraattt gattgttact cattttgctc tcatgtkct	a 539													
aaacagcaac agtgtaacta gtcttttgtt gtaaatggtt attttcctta taaaaattt	t 599													
aaaaactaaq tqqcaaattc catgaaaata tttctcagtt ctgtatgcac ttttattta	a 659													
cattattoat ataattotoo coccaccact ttatttataa ataotgoaaa aktgaraag	g 719													
agataataaa tactttgoto tgaatttggo atocaaagtt aacatttoto cootcacto	c 779													
cttqctqqtq tcataqttat tagaatcagc agcctcttaa ctaattgcgg tttcatagg	a 839													
tatataaatg tttcaagcca ttattgctga atggttcttt agttattaac ctagaccca	a 899													
atcaaagacc agttggattt atgatatttt ttatttgttc ttgcagccaa agtgccagt	t 959 t 1019													
totttaatat gtgaccaaga acacaaggag catcoatatg gocaaataaa tacactgaa	1019													
tttagaaaaa caaaaaaaaa ar	1041													

<210> 374 <211> 1164 <212> DNA <213> Homo sapiens <220>

<222> 72..545

<221> CDS

<221> sig_peptide <222> 72..203

<223> Von Heijne matrix score 5.5 seq ILFFTGWWIMIDA/AV

<221> polyA_site <222> 1151..1162

	-30					-25					-20					
Ala -15	ggt Gly	Ile	Leu	Phe	Phe	Thr	Gly	Trp	Trp	Ile	Met	Ile	Asp	gca Ala	Ala 1	206
ata	gtg Val	tat Tyr	cct Pro 5	aag Lys	cca Pro	gaa Glu	cag Gln	ttg Leu 10	aac Asn	cat His	gcc Ala	ttt Phe	cac His 15	aca Thr	tgt Cys	254
ggt Gly	gta Val	ttt Phe 20	tcc Ser	aca Thr	ttg Leu	gct Ala	ttc Phe 25	ttc Phe	atg Met	ata Ile	aat Asn	gct Ala 30	gta Val	tcc Ser	aat Asn	302
gct Ala	cag Gln 35	ata	aga Arg	ggt Gly	gat Asp	agc Ser 40	tat Tyr	gaa Glu	agc Ser	ggc Gly	tgt Cys 45	tta Leu	gga Gly	aga Arg	aca Thr	350
ggt Gly 50	act	cga Arg	gtt Val	tgg Trp	ctt Leu 55	ttc Phe	att Ile	ggt Gly	ttc Phe	atg Met 60	ttg Leu	atg Met	ttt Phe	gjà aaa	tca Ser 65	398
ctt	att Ile	gct Ala	\tcc Ser	atg Met 70	tgg	att Ile	ctt Leu	ttt Phe	ggt Gly 75	gca Ala	tat Tyr	gtt Val	acc Thr	caa Gln 80	aat Asn	446
act Thr	gat Asp	gtt Val	tat Tyr 85	ccq	gga Gly	cta Leu	gct Ala	gtg Val 90	ttt Phe	ttt Phe	caa Gln	aat Asn	gca Ala 95	ctt Leu	ata Ile	494
ttt Phe	ttt Phe	agc Ser 100	act	ctg Leu	atc Ile	tac Tyr	aaa Lys 105	ttt Phe	gga Gly	aga Arg	acc Thr	gaa Glu 110	gag Glu	cta Leu	tgg Trp	542
	tga		act	tctt	aagt	ca c	attt	tcct	t tt	gtta	tatt	ctg	tttg	tag		595
Thr ataggtttt tatctctag tacacattge casatggagt agattgtaca ttaaatgttt tgtttctta catttttatg ttctgagttt tgaaatagtt ttatgaaatt tctttattt tcattgcata gactgttaat atgtatataa tacaagacta tatgaattgg ataatgagta tcagttttt tatcctgaga tttagaactt gatctactce ctgagccagg gttacatcat cttgtcatt tagaagtaac cactcttgte tctctggctg ggcacggtgg ctcatgcctg taatcccage actttgggag gccgaggcgg gccgattgct tgaggtcaag tgtttgagac cagcctggcc aacatggcga aaccccatct actaaaaata caaaaattag ccaggcatgg tggtgggtgc ctgtaatccc aactacctag gaggctgagg caggagaatc gcttgaaccc ggggggcaga ggttgyagtg agctgagttt gcgccactge actctagcct ggggggagaaa gtgaaactcc ctctcaaaaa aaaaaaamc													655 715 775 835 895 955 1015 1075 1135			

```
<210> 375
```

<211> 1250

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 36..425

<221> sig_peptide

<222> 36..119

<223> Von Heijne matrix score 11.6000003814697 seq LLLLVQLLRFLRA/DG

<221> polyA_signal <222> 1215..1220

<221> polyA_site <222> 1240..1250

<400> 375

atttcttccc cccgagctgg gcgtgcgcgg ccgca atg aac tgg gag ctg ctg Met Asn Trp Glu Leu -25	53
ctg tgg ctg ctg gtg ctg gcg ctg ctc ctg ctc ttg gtg cag ctg Leu Trp Leu Leu Val Leu Cys Ala Leu Leu Leu Leu Val Gln Leu -20 -15 -10	101
ctg cgc ttc ctg agg gct gac ggc gac ctg acg cta cta tgg gcc gag Leu Arg Phe Leu Arg Ala Asp Gly Asp Leu Thr Leu Leu Trp Ala Glu -5 10	149
tgg cag gga cga cgc cca gaa tgg gag ctg act gat atg gtg gtg tgg Trp Gln Gly Arg Arg Pro Glu Trp Glu Leu Thr Asp Met Val Val Trp 15 20 25	197
gtg act gga gcc tcg agt gga att ggt gag gag ctg gct tac cag ttg Val Thr Gly Ala Ser Ser Gly Ile Gly Glu Glu Leu Ala Tyr Gln Leu 30 35 40	245
tct aaa cta gga gtt tct ctt gtg ctg tca gcc aga aga gtg cat gag Ser Lys Leu Gly Val Ser Leu Val Leu Ser Ala Arg Arg Val His Glu 45 50 55	293
ctg gaa agg gtg aaa aga aga tgc cta gag aat ggc aat tta aaa gaa Leu Glu Arg Val Lys Arg Arg Cys Leu Glu Asn Gly Asn Leu Lys Glu 60 65 70	341
aaa gat ata ctt gtt ttg ccc ctt gac ctg acc gac act ggt tcc cat Lys Asp Ile Leu Val Leu Pro Leu Asp Leu Thr Asp Thr Gly Ser His 75 80 85 90	389
gaa agc ggc tac caa agc tgt tct cca gga att tgg tagaatcgac Glu Ser Gly Tyr Gln Ser Cys Ser Pro Gly Ile Trp 95 100	435
attotggtca acaatgtgga aatgtcccag cgttctctgt gcatggatac caacttggat	495
gtctacagaa agctaatgag agcttaacta cttagggacg gtgtccttga caaaatgtgk	555
kctgcctcac atgatcgaga ngaarcaagg aaagattgtt actgtgaata gcatcctggg	615
tatcatatot gracetetti ceattggata etgigetage aageatgete teeggyykik	675
ktttaatggc cttcraacag aacttgccac atacccargt ataatagttt ctaacatttg	735 795
cccaggacct gtgcaatcaa atattgtgga aaattcccta gctggagaag tcacaaagac	855
tataggcaat aatggagacc agtcccacaa gatgacaacc agtcgttgtg tgcggctgat	915
gttaatcagc atggccaatg atttgaaaga agtttggatc tcagaacaac ctttcttgtt	975
agtaacatat ttgtggcaat acatgccaac ctgggcctgg tggataacca acaagatggg	1035
gaagaaaagg attgagaact ttaagagtgg tgtggatgca gactcttctt attttaaaat ctttaagaca aaacatgact gaaaagagca cctgtacttt tcaagccact ggagggagaa	1095
atggaaaaca tgaaaacagc aatcttctta tgcttctgaa taatcaaaga ctaattgtg	1155
attttacttt ttaatagata tgactttgct tccaacatgg aatgaaataa aaaataaata	1215
ataaaagatt gccatgaatc ttgcaaaaaa aaaaa	1250

<210> 376

<211> 947

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 155..751

<221> sig_peptide

<222> 155..340 <223> Von Heijne matrix score 3.70000004768372 seq SILGIISVPLSIG/YC

<221> polyA_signal <222> 912..917

<221> polyA_site <222> 937..947

<400	> 37	6											+		++===	60
agtg	aaaa	ga a	gatg	ccta	g ag	aatg	gcaa	ttt	aaaa	gaa	aaag	atat	ac t	eget	ttgcc	120
cctt	dacc	to a	ccga	cact	a at	tece	atqa	ago	ggct	acc	aaag	ctgt	. בכי ב	.cca9	gagee	175
tggt	agaa	tc g	jacat	tctg	g tc	aaca	atgg	tgg	a at	gto	c ca	g cg	וכ נכ	ינ נינ	.g tgc	1/3
									Me	t Se	r G1	n Ar	g se	i re	u Cys	
											-6	_			4	223
atg	gat	acc	agc	ttg	gat	gtc	tac	aga	rag	cta	ata	gag	CEE	aac	Tac	223
Met	Asp	Thr	Ser	Leu	Asp	Val	Tyr	Arg	Xaa	Leu	Ile	Glu	Leu	Asn.	TAL	
-55					-50					-45					-40	071
tta	ggg	acg	gtg	tcc	ttg	aca	aaa	tgt	gtt	ctg	cct	cac	atg	atc	gag	271
Leu	Gly	Thr	Val	Ser	Leu	Thr	Lys	Cys	Val	Leu	Pro	His	Met	TIE	Glu	
				-35					-30					-25		
agg	aaq	caa	qqa	aag	att	gtt	act	gtg	aat	agc	atc	ctg	ggt	atc	ata	319
Ara	Lvs	Gln	Gly	Lys	Ile	Val	Thr	Val	Asn	Ser	Ile	Leu	Gly	Ile	Ile	
			-20					-15					-10			
tct	gta	cct	ctt	tcc	att	qqa	tac	tgt	gct	agc	aag	cat	gct	ctc	cgg	367
Ser	Val	Pro	Leu	Ser	Ile	Glv	Tyr	Cys	Ala	Ser	Lys	His	Ala	Leu	Arg	
501		-5				•	1	-			5					
aat		+++	aat	aac	ctt	caa	aca	gaa	ctt	gcc	aca	tac	cca	ggt	ata	415
330	Dhe	Dhe	Asn	GIV	Leu	Ara	Thr	Glu	Leu	Ala	Thr	Tyr	Pro	Gly	Ile	
10	1110	1110	11011	017	15	5				20					25	
10	att	tct	aac	att	tac	cca	aga	cct	ata	caa	tca	aat	att	gtg	gaa	463
Tla	Val	Ser	Asn	Tle	Cvs	Pro	Glv	Pro	Val	Gln	Ser	Asn	Ile	Val	Glu	
116	vai	DCI	ADII	30	- 22	• • •	2		35					40		
22t	tcc	cta	act	aga	gaa	atc	aca	aaa	act	ata	ggc	aat	aat	gga	aac	511
A m m	Sar	Len	Z la	Glv	Glu	Val	Thr	Lvs	Thr	Ile	Gly	Asn	Asn	Cly	Asn	
14511	~ C 4.	عا جائد	45	OI,	014			50					55			
626	tcc	Cac	220	ato	aca	acc	agt		tat	ata	cqq	ctg	atg	tta	atc	559
cay	COX	ui.c	Tyc	Mat	Thr	Thr	Ser	Ara	Cvs	Val	Arg	Leu	Met	Leu	Ile	
GIII	261	60	دلاند	Mec	1111	****	65	3	-1-		5	70				
		~~~	<del>+</del>	ant-	++~	222		att	taa	atc	tca	gaa	caa	cct	ttc	607
agc	atg	310	aac Nen	yac Nan	Leu	Luc	Glu	Val	Tro	Tle	Ser	Glu	Gln	Pro	Phe	
ser		Ald	ASII	Asp	Бец	80	014	,			85					
***	75	~+ -	202	+ - +	++~		caa	tac	ato	cca		taa	qcc	tgg	tgg	655
ttg	Tan	yea	The	Tree	Leu	Trn	Gln	Tyr	Met	Pro	Thr	Trp	Āla	Trp	Trp	
	ьeu	Val	1111	TYL	95	115	0111	- 7 -		100				•	105	
90					95	224	222	200	att		aac	ttt	aad	agt	aat	703
ata	acc	aac	aag	atg	999	aay	T	299	Tla	Glu	Acn	Dhe	Lvs	Ser	Glv	
Ile	Thr	Asn	гуs		GIY	гуs	пув	Arg	115	GIU	no	1	_,_	120	2	
				110							220	202	222		gac	751
gtg	gat	gcm	rac	tct	ECE	tat	בענ	aaa	TIO	Dho	Lag	Thr	Live	cat	Asp	
Val	Asp	Ala			Ser	туг	Pne	гуs	116	Pne	пуъ	1111	135	His		
			125	,			~	130	<b>~</b> ~ ~ ~	~~~	27+				aaacad	811
tga	aaag	anc	acct	gtac	tt t	tcaa	gcca	c tg	yayy	yaya	aat	yyaa +++>	aac a++	+++=	aaacag	871
caa	tctt	ctt	atgc	ttct	ga a	caat	caaa	g ac	taat	ctgt	gat	2222	~ T	+000	atagat	931
					tg g	rrtg	aaat	a aa	aaat	aaat	aat	aadd	yaı	Lyce	atgrrt	947
ctt	gcaa	aaa	aaaa	.aa												

<210> 377

<211> 621

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 46..585

<221> sig_peptide

621

<222> 46..120 <223> Von Heijne matrix score 6.30000019073486 seq AFSLSVMAALTFG/CF <221> polyA_signal <222> 584..589 <221> polyA_site <222> 606..619 <400> 377 aactgggtgt gcgtrtggag tccggactcg tgggagacga tcgcg atg aac acg gtg 57 Met Asn Thr Val 105 ctg tcg cgg gcg aac tca ctg ttc gcc ttc tcg ctg agc gtg atg gcs Leu Ser Arg Ala Asn Ser Leu Phe Ala Phe Ser Leu Ser Val Met Ala -15 geg etc ace tte gge tge tte atc ayy ace gee tte aaa gae agg age 153 Ala Leu Thr Phe Gly Cys Phe Ile Xaa Thr Ala Phe Lys Asp Arg Ser 1 gtc ccg gtg cgg ctg cac gtc tcg cga atc atg cta aaa aat gta gaa 201 Val Pro Val Arg Leu His Val Ser Arg Ile Met Leu Lys Asn Val Glu 20 15 gat tto act gga cot aga gaa aga agt gat otg gga ttt ato aca ttt Asp Phe Thr Gly Pro Arg Glu Arg Ser Asp Leu Gly Phe Ile Thr Phe 35 297 gat ata act gct gat cta gag aat ata ttt gat tgg aat gtt aag cag Asp Ile Thr Ala Asp Leu Glu Asn Ile Phe Asp Trp Asn Val Lys Gln 55 50 ttg ttt ctt tat tta tca gca gaa tat tca aca aaa aat aat gct ctg 345 Leu Phe Leu Tyr Leu Ser Ala Glu Tyr Ser Thr Lys Asn Asn Ala Leu 70 65 393 aac caa ktt gtc cta tgg gac aag att gtt ttg aga ggt gat aat ccg Asn Gln Xaa Val Leu Trp Asp Lys Ile Val Leu Arg Gly Asp Asn Pro 85 aag ctg ctg ctg aaa gat atg aaa aca aaa tat ttt ttc ttt gac gat 441 Lys Leu Leu Lys Asp Met Lys Thr Lys Tyr Phe Phe Asp Asp 100 95 gga aat ggt ctc wag gga aac agg aat gtc act ttg acc ctg tct tgg 489 Gly Asn Gly Leu Xaa Gly Asn Arg Asn Val Thr Leu Thr Leu Ser Trp 115 110 537 aac gtc gta cca aat gct gga att cta cct ctt gtg aca gga tca gga Asn Val Val Pro Asn Ala Gly Ile Leu Pro Leu Val Thr Gly Ser Gly

130

145

taaattatto tgaatttgaa acaaaaaaa aaaahm

cac gta tct gtc cca ttt cca gat aca tat gaa ata acg aag agt tat His Val Ser Val Pro Phe Pro Asp Thr Tyr Glu Ile Thr Lys Ser Tyr

135

<210> 378

<211> 52

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -20..-1

<400> 378

<210> 379 <211> 193 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -23..-1 <400> 379 Met Val Val Leu Arg Ala Gly Lys Lys Thr Phe Leu Pro Pro Leu Xaa -15 -20 Arg Ala Phe Ala Cys Arg Gly Cys Gln Leu Ala Pro Glu Arg Gly Ala Glu Arg Arg Asp Thr Ala Pro Ser Gly Val Ser Arg Phe Cys Pro Pro 20 15 Arg Lys Ser Cys His Asp Trp Ile Gly Pro Pro Asp Lys Tyr Ser Asn 35 3.0 Leu Arg Pro Val His Phe Tyr Ile Pro Glu Asn Glu Ser Pro Leu Glu 50 Gln Lys Leu Arg Lys Leu Arg Gln Glu Thr Gln Glu Trp Asn Gln Gln 65 Phe Trp Ala Asn Gln Asn Leu Thr Phe Ser Lys Glu Lys Glu Glu Phe 80 Ile His Ser Arg Leu Lys Thr Lys Gly Leu Gly Leu Arg Thr Glu Ser 100 95 Gly Gln Lys Ala Thr Leu Asn Ala Glu Glu Met Ala Asp Phe Tyr Lys 115 110 Glu Phe Leu Ser Lys Asn Phe Gln Lys His Met Tyr Tyr Asn Arg Asp 125 130 Trp Tyr Lys Arg Asn Phe Ala Ile Thr Phe Phe Met Gly Lys Val Ala 150 145 Leu Glu Arg Ile Trp Asn Lys Leu Lys Gln Lys Gln Lys Lys Arg Ser 160 Asn

<210> 380 <211> 82 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1

170

<210> 381 <211> 198 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 381 Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala Leu Ala Met Val Thr -10 -15 Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro Glu Leu Ala Gln His <del>-</del> 5 Glu Glu Leu Thr Leu Leu Phe His Gly Thr Leu Gln Leu Gly Gln Ala 20 15 Leu Asn Gly Val Tyr Arg Thi Thr Glu Gly Arg Leu Thr Lys Ala Arg 35 Asn Ser Leu Gly Leu Tyr Gly Arg Thr Ile Glu Leu Leu Gly Gln Glu 50 Val Ser Arg Gly Arg Asp Ala Ala Gln Glu Leu Arg Ala Ser Leu Leu 70 65 Glu Thr Gln Met Glu Glu Asp Ile Leu Xaa Leu Gln Ala Xaa Ala Thr 85 Ala Glu Val Leu Gly Glu Val Ala Gln Ala Gln Lys Val Leu Arg Asp 100 95 Ser Val Gln Arg Leu Xaa Xaa Gln Leu Xaa Xaa Ala Trp Leu Gly Pro 120 115 110 Ala Tyr Arg Lys Phe Glu Val Leu Lys Ala Pro Pro Xaa Lys Gln Asn 135 130 125 His Ile Leu Trp Ala Leu Thr Gly His Val Xaa Arg Gln Xaa Arg Glu 150 145 140 Met Val Ala Gln Gln Xaa Xaa Leu Xaa Gln Ile Gln Glu Lys Leu His 165 160 Thr Ala Ala Leu Pro Ala 175

<210> 382 <211> 160 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -55..-1

Phe Gly

<400> 382 Met Asp Lys Leu Lys Lys Val Leu Ser Gly Gln Asp Thr Glu Asp Arg

-45 -50 Ser Gly Leu Ser Glu Val Val Glu Ala Ser Ser Leu Ser Trp Ser Thr -25 · -30 -35 Arg Ile Lys Gly Phe Ile Ala Cys Phe Ala Ile Gly Ile Leu Cys Ser -15 -20 Leu Leu Gly Thr Val Leu Leu Trp Val Pro Arg Lys Gly Leu His Leu 1 Phe Ala Val Phe Tyr Thr Phe Gly Asn Ile Ala Ser Ile Gly Ser Thr 20 15 Ile Phe Leu Met Gly Pro Val Lys Gln Leu Lys Arg Met Phe Glu Pro 35 30 Thr Arg Leu Ile Ala Thr Ile Met Val Leu Leu Cys Phe Ala Leu Thr 50 45 Leu Cys Ser Ala Phe Trp Trp His Asn Lys Gly Leu Ala Leu Ile Phe 65 Cys Ile Leu Gln Ser Leu Ala Leu Thr Trp Tyr Ser Leu Ser Phe Ile 80 Pro Phe Ala Arg Asp Ala Val Lys Xaa Cys Phe Ala Val Cys Leu Ala 100

<210> 383 <211> 108 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -18..-1

<210> 384 <211> 64 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1

Leu Tyr Ile Pro Xaa Arg Xaa Arg Ser Asp Glu Leu Val Phe Glu Ser 15 20 25 .

Gln Lys Gly Ser Ala Met Glu Leu Ala Val Ile Thr Val Xaa Gly Val 30 35

<210> 385

<210> 386 <211> 186 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 386 Met Ser Pro Ser Gly Arg Leu Cys Leu Leu Thr Ile Val Gly Leu Ile -10 -15 Leu Pro Thr Arg Gly Gln Thr Leu Lys Asp Thr Thr Ser Ser Ser Ser 1 Ala Asp Ser Thr Ile Met Asp Ile Gln Val Pro Thr Arg Ala Pro Asp 20 15 Ala Val Tyr Thr Glu Leu Gln Pro Thr Ser Pro Thr Pro Thr Trp Pro 35 30 Ala Asp Glu Thr Pro Gln Pro Gln Thr Gln Thr Gln Gln Leu Glu Gly 50 Thr Asp Gly Pro Leu Val Thr Asp Pro Glu Thr His Xaa Ser Xaa Lys Ala Ala His Pro Thr Asp Asp Thr Thr Leu Ser Glu Arg Pro Ser 85 Pro Ser Thr Xaa Val His Xaa Arg Pro Xaa Xaa Pro Ser Xaa His Leu 100 Val Phe Met Arg Met Thr Pro Ser Ser Met Met Asn Thr Pro Ser Gly 120 115 Asn Xaa Gly Cys Trp Ser Gln Leu Cys Cys Ser Ser Gln Ala Ser Ser 135 130 Ser Ser Pro Val Ala Ser Ala Gly Ser Cys Pro Gly Tyr Ala Gly Ile 150 145 Ile Ala Gly Glu Ser Ile Arg Asn Arg Ser

```
<210> 387
<211> 179
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -26..-1
<400> 387
Met Glu Thr Gly Ala Leu Arg Arg Pro Gln Leu Leu Pro Leu Leu
                                 -15
                       -20
Leu Cys Gly Pro Ser Gln Asp Gln Cys Arg Pro Val Leu Gln Asn
                   -5
Leu Leu Gln Ser Pro Gly Leu Thr Trp Ser Leu Glu Val Pro Thr Gly
                              15
           10
Arg Glu Gly Lys Glu Gly Gly Asp Arg Gly Pro Gly Leu Xaa Gly Ala
                           30
       25
Thr Pro Ala Arg Ser Pro Gln Gly Lys Glu Met Gly Arg Gln Arg Thr
                       45
Arg Lys Val Lys Gly Pro Ala Trp Xaa His Thr Ala Asn Gln Glu Leu
                  60
Asn Arg Met Arg Ser Leu Ser Ser Gly Ser Val Pro Val Gly His Leu
Glu Gly Gly Thr Val Lys Leu Gln Lys Asp Thr Gly Leu His Ser Cys
                                95
           90
Xaa Asp Gly Met Ala Ser Leu Glu Gly Thr Pro Ala Ser Val Leu Ala
                                              115
                            110
Asp Ala Cys Pro Gly Phe His Asp Val Xaa Val Gln Xaa Ala Leu Phe
                                           130
                        125
Gly Leu Ser Gly Xaa Xaa Leu Trp Leu Lys Thr His Phe Cys Leu Ser
                                       145
               140
 135
 Ile Xaa Leu
 <210> 388
 <211> 150
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -55..-1
 Met Ala Thr Thr Val Pro Asp Gly Cys Arg Asn Gly Leu Lys Ser Lys
                                        -45
                    -50
 Tyr Tyr Arg Leu Cys Asp Lys Ala Glu Ala Trp Gly Ile Val Leu Glu
                                    -30
                 -35
 Thr Val Ala Thr Ala Gly Val Val Thr Ser Val Ala Phe Met Leu Thr
                                 -15
             -20
 Leu Pro Ile Leu Val Cys Lys Val Gln Asp Ser Asn Arg Arg Lys Met
         -5
 Leu Pro Thr Gln Phe Leu Phe Leu Leu Gly Val Leu Gly Ile Phe Gly
                                        20
                     15
 10
 Leu Thr Phe Ala Phe Ile Ile Gly Leu Asp Gly Ser Thr Gly Pro Thr
                                    35
                 30
 Arg Phe Phe Leu Phe Gly Ile Leu Phe Ser Ile Cys Phe Ser Cys Leu
                                50
             45
 Leu Ala His Ala Val Ser Leu Thr Lys Leu Val Arg Gly Arg Lys Ala
                                                 70
                           65
```

Pro Phe Pro Val Gly Asp Ser Gly Ser Gly Arg Gly Leu Gln Pro Ser 75 80 85

Pro Gly Cys Tyr Arg Tyr 90 95

<210> 389 <211> 236 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -31..-1 <400> 389 Met Leu Ser Lys Gly Leu Lys Arg Lys Arg Glu Glu Glu Glu Lys -20 -25 Glu Pro Leu Ala Val Asp Ser Trp Trp Leu Asp Pro Gly His Ala Ala - 5 -10 Val Ala Gln Ala Pro Pro Ala Val Ala Ser Ser Leu Phe Asp Leu 10 Ser Val Leu Lys Leu His His Ser Leu Gln Xaa Ser Xaa Pro Asp Leu 25 Arg His Leu Val Leu Val Xaa Asn Thr Leu Arg Arg Ile Gln Ala Ser 45 40 Met Ala Pro Ala Ala Leu Pro Pro Val Pro Thr Pro Pro Ala Ala 60 Pro Xaa Val Ala Asp Asn Leu Leu Ala Ser Ser Asp Ala Ala Leu Ser 75 70 Ala Ser Met Ala Xaa Leu Leu Glu Asp Leu Ser His Ile Glu Gly Leu 90 Ser Gln Ala Pro Gln Pro Leu Ala Asp Glu Gly Pro Pro Gly Arg Ser 110 105 Ile Gly Gly Xaa Pro Pro Xaa Leu Gly Ala Leu Asp Leu Leu Gly Pro 120 125 Ala Thr Gly Cys Leu Leu Asp Asn Gly Leu Glu Gly Leu Phe Glu Asp 140 135 Ile Asp Thr Ser Met Tyr Asp Asn Glu Leu Trp Ala Pro Ala Ser Glu 155 150 Gly Leu Lys Pro Gly Pro Glu Asp Gly Pro Gly Lys Glu Glu Ala Pro 175 170 165 Glu Leu Asp Glu Ala Glu Leu Asp Tyr Leu Met Asp Val Leu Val Gly 185 Thr Gln Ala Leu Glu Arg Pro Pro Gly Pro Gly Arg 200

<210> 390 <211> 149 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -100..-1

<400> 390
Met Glu Thr Leu Tyr Arg Val Pro Phe Leu Val Leu Glu Cys Pro Asn
-100 -95 -90 -85

Leu Lys Leu Lys Lys Pro Pro Trp Leu His Met Pro Ser Ala Met Thr -75 -70 -80 Val Tyr Ala Leu Val Val Val Ser Tyr Phe Leu Ile Thr Gly Gly Ile -60 Ile Tyr Asp Val Ile Val Glu Pro Pro Ser Val Gly Ser Met Thr Asp -45 Glu His Gly His Gln Arg Pro Val Ala Phe Leu Ala Tyr Arg Val Asn -30 Gly Gln Tyr Ile Met Glu Gly Leu Ala Ser Ser Phe Leu Phe Thr Met -15 -10 Gly Gly Leu Gly Phe Ile Ile Leu Asp Gly Ser Asn Ala Pro Asn Ile 1 Pro Lys Leu Asn Arg Phe Leu Leu Leu Phe Ile Gly Phe Val Cys Val 20 Leu Xaa Ser Phe Xaa Xaa Ala Arg Val Phe Met Arg Met Lys Leu Pro Gly Tyr Leu Met Gly

<210> 391
<211> 69
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -49..-1

<210> 392 <211> 241 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1

```
Cys Leu Trp Phe Arg Tyr Gly Ala His Gln Pro Glu Asn Leu Cys Leu
                  40
                                       45
Asp Gly Cys Lys Ser Glu Ala Xaa Lys Phe Thr Val Arg Glu Ala Leu
                                   60
               55
Lys Glu Asn Gln Val Ser Leu Thr Val Asn Arg Val Thr Ser Asn Asp
Ser Ala Ile Tyr Ile Cys Gly Ile Ala Phe Pro Ser Val Pro Glu Ala
                           90
Arg Ala Lys Gln Thr Gly Gly Gly Thr Thr Leu Val Val Arg Glu Ile
                       105
                                           110
Lys Leu Leu Ser Lys Glu Leu Arg Ser Phe Leu Thr Ala Leu Val Ser
                                      125
                   120
Leu Leu Ser Val Tyr Val Thr Gly Val Cys Val Ala Phe Ile Leu Leu
                                   140
               135
Ser Lys Ser Lys Ser Asn Pro Leu Arg Asn Lys Glu Ile Lys Glu Asp
                              155
Ser Gln Lys Lys Ser Ala Arg Arg Ile Phe Gln Glu Ile Ala Gln
                                        175
                          170
Glu Leu Tyr His Lys Arg His Val Glu Thr Asn Gln Gln Ser Glu Lys
                      185
                                       190
Asp Asn Asn Thr Tyr Glu Asn Arg Arg Val Leu Ser Asn Tyr Glu Arg
                  200
Pro
```

<210> 393 <211> 47 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1

<210> 394 <211> 65 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1

Ser

<210> 395

<211> 73 <212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -24..-1

<400> 395

Met Thr Cys Trp Met Leu Pro Pro Ile Ser Phe Leu Ser Tyr Leu Pro -20

-15

Leu Trp Leu Gly Pro Ile Trp Pro Cys Ser Gly Ser Thr Leu Gly Lys 1

Pro Asp Pro Gly Val Trp Pro Ser Leu Phe Arg Pro Trp Asp Ala Ala

15

Ser Pro Gly Asn Tyr Ala Leu Ser Arg Gly Xaa Asn Xaa Tyr Xaa Xaa

30

Trp Gly Gln Gly Thr His Ser Ser Leu

45

<210> 396

<211> 60

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -18..-1

<400> 396

Met Pro Cys Pro Thr Trp Thr Cys Leu Lys Ser Phe Pro Ser Pro Thr -15 -10 -5

Ser Ser His Ala Ser Ser Leu His Leu Pro Pro Ser Cys Thr Arg Leu

10 Thr Leu Thr Gln Thr Leu Arg Thr Gly Met His Leu Ser Arg Ala Leu

20 25

Gln Gly Thr Leu Thr Arg Leu Gln Ser Thr Pro Ala 35

<210> 397

<211> 192

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -93..-1

<400> 397

Met Ala Glu Leu Gly Leu Asn Glu His His Gln Asn Glu Val Ile Asn - 90 -85

Tyr Met Arg Phe Ala Arg Ser Lys Arg Gly Leu Arg Leu Lys Thr Val

-70 -65 Asp Ser Cys Phe Gln Asp Leu Lys Glu Ser Arg Leu Val Glu Asp Thr -50 -55 Phe Thr Ile Asp Glu Val Ser Glu Val Leu Asn Gly Leu Gln Ala Val -35 -40 Val His Ser Glu Val Glu Ser Glu Leu Ile Asn Thr Ala Tyr Thr Asn -20 Val Leu Leu Leu Arg Gln Leu Phe Ala Gln Ala Glu Lys Trp Tyr Leu -5 Lys Leu Gln Thr Asp Ile Ser Glu Leu Glu Asn Arg Glu Leu Leu Glu 10 15 Gln Xaa Ala Glu Phe Glu Lys Ala Xaa Ile Thr Ser Ser Asn Lys Lys 30 25 Pro Ile Leu Xaa Val Thr Xaa Pro Lys Leu Ala Pro Leu Asn Glu Gly 45 40 Gly Thr Ala Lys Leu Leu Asn Lys Val Ile Cys Ile Ile Leu Arg Asn 60 √55 Gly Lys Ser Leu Ile Leu Ser Cys His Cys Leu Gly Trp Arg Asn Lys 80 75 Ser Gly Arg Phe Val Ser Gly Pro Leu Arg Ile Ile Ser Pro Leu Gln 90

<210> 398 <211> 149 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -72..-1

<400> 398 Met Asn Leu Phe Ile Met Tyr Met Ala Gly Asn Thr Ile Ser Ile Phe -65 -60 Pro Thr Met Met Val Cys Met Met Ala Trp Arg Pro Ile Gln Ala Leu -50 Met Ala Ile Ser Ala Thr Phe Lys Met Leu Glu Ser Ser Ser Gln Lys -30 -35 Phe Leu Gln Gly Leu Val Tyr Leu Ile Gly Asn Leu Met Gly Leu Ala -15 -20 Leu Ala Val Tyr Lys Cys Gln Ser Met Gly Leu Leu Pro Thr His Ala 1 <del>-</del>5 Ser Asp Trp Leu Ala Phe Ile Glu Pro Pro Glu Arg Met Glu Ser Val 15 Val Glu Asp Cys Phe Cys Glu His Glu Lys Ala Ala Pro Gly Pro Tyr 30 35 Val Phe Gly Ser Tyr Leu His Pro Ser Leu Ser Pro Val Ala Pro Gln 50 4.5 His Thr Leu Lys Leu Ile Thr Tyr Val Lys Lys Asn Gln Lys Thr Leu 60 Phe Ser Met Val Gly

<210> 399 <211> 73 <212> PRT <213> Homo sapiens

<220>
<221> SIGNAL
<222> -20..-1

<210> 400 <211> 86 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -20..-1

<400> 400 Met Asn Leu His Phe Pro Gln Trp Phe Val His Ser Ser Ala Leu Gly -10 -15 Leu Val Leu Ala Pro Pro Phe Ser Ser Pro Gly Thr Asp Pro Thr Phe 1 Pro Cys Ile Tyr Cys Arg Leu Leu Asn Met Ile Met Thr Arg Leu Ala 20 1.5 Phe Ser Phe Ile Thr Cys Leu Cys Pro Asn Leu Lys Glu Val Cys Leu 40 35 Ile Leu Pro Glu Lys Asn Cys Asn Ser Arg His Ala Gly Phe Val Gly 55 50 Pro Xaa Lys Leu Arg Gln 65

50

55

<210> 402 <211> 65 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1

<210> 403 <211> 211 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1

<400> 403 Met Leu Leu Ser Ile Thr Thr Ala Tyr Thr Gly Leu Glu Leu Thr -20 Phe Phe Ser Gly Val Tyr Gly Thr Cys Ile Gly Ala Thr Asn Lys Phe - 5 Gly Ala Glu Glu Xaa Ser Leu Ile Gly Leu Ser Gly Ile Phe Ile Gly 10 15 Ile Gly Glu Ile Leu Gly Gly Ser Leu Phe Gly Leu Leu Ser Lys Asn 30 Asn Arg Phe Gly Arg Asn Pro Val Val Leu Leu Gly Ile Leu Val His 45 Phe Ile Ala Phe Tyr Leu Ile Phe Leu Asn Met Pro Gly Asp Ala Pro 60 Ile Ala Pro Val Lys Gly Thr Asp Ser Ser Ala Tyr Ile Lys Ser Ser 75 Lys Xaa Phe Ala Ile Leu Cys Xaa Phe Leu Xaa Gly Leu Gly Asn Ser 95 Cys Phe Asn Thr Xaa Leu Leu Xaa Ile Xaa Gly Phe Leu Tyr Ser Glu 110 Xaa Ser Ala Pro Xaa Phe Ala Ile Phe Asn Phe Val Gln Ser Ile Cys 125 Ala Ala Val Ala Phe Phe Tyr Ser Asn Tyr Leu Leu Leu His Trp Gln 140 145 Leu Leu Val Met Val Ile Phe Gly Phe Xaa Gly Thr Ile Ser Phe Phe 150 - 155 160 Thr Val Glu Trp Glu Xaa Ala Ala Phe Val Xaa Arg Gly Ser Asp Tyr 170

Arg Ser Ile

<210> 404 <211> 123 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -80..-1 <400> 404

Met Ser Thr Trp Tyr Leu Ala Leu Asn Lys Ser Tyr Lys Asn Lys Asp -70 -75 Ser Val Arg Ile Tyr Leu Ser Leu Cys Thr Val Ser Ile Lys Phe Thr -55 -60 Tyr Phe His Asp Ile Gln Thr Asn Cys Leu Thr Thr Trp Lys His Ser -40 -45 Arg Cys Arg Phe Tyr Trp Ala Phe Gly Gly Ser Ile Leu Gln His Ser -25 -20 -30 Val Asp Pro Leu Val Leu Phe Leu Ser Leu Ala Leu Leu Val Thr Pro -15 -10 Thr Ser Thr Pro Ser Ala Lys Ile Gln Ser Leu Gln Ile Asp Leu Pro 5 10 Gly Gly Trp Arg Leu Ala Thr Asp Arg Ile Phe Thr Leu Ser Pro Val 25 20 Pro Met Asp Xaa Pro Leu Ile Leu His Gln Leu

40

<210> 405 <211> 86 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1

<400> 405 Met Glu Lys Ser Trp Met Leu Trp Asn Phe Val Glu Arg Trp Leu Ile -20 -15 Ala Leu Ala Ser Trp Ser Trp Ala Leu Cys Arg Ile Ser Leu Leu Pro 1 - 5 Leu Ile Val Thr Phe His Leu Tyr Gly Gly Ile Ile Leu Leu Leu 10 15 Ile Phe Ile Ser Ile Xaa Gly Ile Leu Tyr Lys Phe Xaa Asp Val Leu 30 Leu Tyr Phe Pro Xaa Gln Xaa Ser Ser Ser Arg Leu Tyr Asp Ser His 50 Ala His Trp Xaa Ser Xaa 55

<210> 406 <211> 162 <212> PRT <213> Homo sapiens

```
<220>
 <221> SIGNAL
  <222> -31..-1
  <400> 406
 Met Ala Ala Arp Pro Ser Gly Pro Xaa Ala Pro Glu Ala Val Thr
                                 -20
                      -25
 Ala Arg Leu Val Gly Val Leu Trp Phe Val Ser Val Thr Thr Gly Pro
                    -10
                                        - 5
 Trp Gly Ala Val Ala Thr Ser Ala Gly Gly Glu Glu Ser Leu Lys Cys
                                10
 Glu Asp Leu Lys Val Gly Gln Tyr Ile Cys Lys Asp Pro Lys Ile Asn
                             25
  Asp Ala Thr Gln Glu Pro Val Asn Cys Thr Asn Tyr Thr Ala His Val
                        40
  Ser Cys Phe Pro Ala Pro Asn Ile Thr Cys Lys Asp Ser Ser Gly Asn
                     55
 Glu Thr His Phe Thr Gly Asn Glu Val Gly Phe Phe Lys Pro Ile Ser
                                     75
. Cys Arg Asn Val Asn Gly Tyr Ser Tyr Asn Glu Gln Ser His Val Ser
                                90
  Phe Ser Trp Met Val Gly Ser Arg Ser Ile Leu Pro Trp Ile Pro Cys
                            105
                                        110
      100
  Phe Gly Phe Val Lys Xaa Xaa His Cys Arg Val Xaa Trp Asn Trp Glu
                     120
  Pro Asn
 130
 <210> 407
  <211> 98
  <212> PRT
  <213> Homo sapiens
 <220>
  <221> SIGNAL
  <222> -37..-1
```

<210> 408 <211> 70 <212> PRT <213> Homo sapiens

PCT/IB98/02122 -

<220> <221> SIGNAL <222> -15..-1 <400> 408 Met Arg Phe Leu Pro Cys Cys Leu Leu Trp Ser Val Phe Asn Pro Glu -15 -10 -5 Ser Leu Asn Cys His Tyr Phe Xaa Xaa Glu Xaa Cys Ile Phe Xaa Ser 15 10 Leu Gln Tyr Tyr Glu Ile Ser Leu Gln Glu Lys Leu Leu Gly Phe Leu - 20 25 Trp Leu Cys Phe Leu Ser Tyr Phe Phe Arg Ala Val Tyr Phe Leu Ile 35 40 Asp Phe Ser Ser Phe Thr 50 55 <210> 409 <211> 60 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -45..-1 <400> 409 Met His Ser Leu Phe Ile Ala Ser Leu Lys Val Leu Phe Tyr Tyr Ser -40 -35 Phe Ser Phe Arg Phe Asn Trp Phe Asp Cys Leu Leu His Asn Leu Gly -25 -20 Glu Asn Phe Leu Ser Leu Leu Ser Lys Ser Cys Ser Ala Asp Pro Ser - 5 -10 Gly Ser Thr Phe Met Arg Asp Ile Glu Thr Asn Lys 10 <210> 410 <211> 39 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1 <400> 410 Met Pro Glu Ala Val Glu Gln Ser Ala His Leu Phe Val Thr Trp Ser -15 -10 Ser Gln Arg Ala Leu Ser His Pro Ala Pro Phe Leu Thr Xaa Xaa Lys 5 10 1

<210> 411 <211> 51 <212> PRT

Asn Pro Phe Leu Trp Lys Leu

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -23..-1

<400> 411

Met Ala Phe Gln Ser Leu Leu Glu Met Lys Phe Phe Leu Cys Ala Ala
-20 -15 -10

Phe Pro Leu Gly Ala Gly Val Lys Met Phe His Tyr Leu Gly Pro Gly
-5 5

Lys Pro Leu Xaa Gln Ala Ser Pro Ser Pro His Pro His Arg Xaa Arg
10 20 25

Ile Trp Pro

<210> 412

<211> 95

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -48..-1

<400> 412

Met Ala Ser Ser His Trp Asn Glu Thr Thr Thr Ser Val Tyr Gln Tyr
-45 -40 -35

Leu Gly Phe Gln Val Gln Lys Ile Tyr Pro Phe His Asp Asn Trp Asn
-30
-25
-20

Thr Ala Cys Phe Val Ile Leu Leu Deu Phe Ile Phe Thr Val Val Ser

~15 -10 -5

Leu Val Val Leu Ala Phe Leu Tyr Glu Val Leu Xaa Xaa Cys Cys Cys 1 5 10 15

Val Lys Asn Lys Thr Val Lys Asp Leu Lys Ser Glu Pro Asn Pro Leu 20 25 30

Xaa Xaa Met Met Asp Asn Ile Arg Lys Arg Glu Thr Glu Val Val 35 40 45

<210> 413

<211> 60

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -32..-1

<400> 413

Met Asp Glu Tyr Ser Trp Trp Cys His Val Leu Glu Val Val Lys Gly
-30 -25 -20

Gln Met Phe Thr Phe Ile Asn Ile Thr Leu Trp Leu Gly Ser Leu Cys

-15 -10 -5
Gln Arg Phe Phe Tyr Ala Ser Gly Thr Tyr Phe Leu Ile Tyr Ile Ser
1 5 10 15

Thr Val Thr Pro Ser Trp Arg Leu Cys Leu Val Ser

```
<210> 414
<211> 170
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -79..-1
<400> 414
Met Glu Asp Pro Asn Pro Glu Glu Asn Met Lys Gln Gln Asp Ser Pro
               -75
                                  -70
Lys Glu Arg Ser Pro Gln Ser Pro Gly Gly Asn Ile Cys His Leu Gly
                               -55
          -60
Ala Pro Lys Cys Thr Arg Cys Leu Ile Thr Phe Ala Asp Ser Lys Phe
                        -40
       -45、
Gln Glu Arg His Met Lys Arg Glu His Pro Ala Asp Phe Val Ala Gln
                                          -20
                       -25
Lys Leu Gln Gly Val Leu Phe Ile Cys Phe Thr Cys Ala Arg Ser Phe
-15 -10
                                       - 5
Pro Ser Ser Lys Ala Xaa Xaa Thr His Gln Arg Ser His Gly Pro Xaa
                                                 15
                               10
Ala Lys Pro Thr Leu Pro Val Ala Thr Thr Thr Ala Gln Pro Thr Phe
                                               30
Pro Cys Pro Asp Cys Gly Lys Thr Phe Gly Gln Ala Val Ser Leu Xaa
Arg His Xaa Gln Xaa His Glu Val Arg Ala Pro Pro Gly Thr Phe Ala
                                      60
Cys Thr Xaa Cys Gly Gln Asp Phe Ala Gln Glu Xaa Gly Leu His Gln
                                  75
              7.0
His Tyr Ile Arg His Ala Arg Gly Gly Leu
           85
<210> 415
<211> 190
<212> PRT
<213> Homo sapiens
```

<210> 415 <211> 190 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -82..-1 <400> 415 Met Tyr Val Trp Pr

Met Tyr Val Trp Pro Cys Ala Val Val Leu Ala Gln Tyr Leu Trp Phe -75 His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly -60 -55 Val Ser Leu Pro Gly Ile Leu Ala Ala Lys Cys Gly Ala Glu Val Ile -45 -40 Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln -20 -25 -30 Ser Cys Gln Met Asn Asn Leu Pro His Leu Gln Val Val Gly Leu Thr -10 Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Xaa Phe Glu Asp Ile 20 25 Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln Leu 35 40

<210> 416 <211> 114 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -60..-1 Met Met Ala Ala Val Pro Pro Gly Leu Glu Pro Trp Asn Arg Val Arg - 55 ~50 Ile Pro Lys Ala Gly Asn Arg Ser Ala Val Thr Val Gln Asn Pro Gly -35 -40 Ala Ala Leu Asp Leu Cys Ile Ala Ala Val Ile Lys Glu Cys His Leu -15 ~20 -25 Val Ile Leu Ser Leu Lys Ser Gln Thr Leu Asp Ala Glu Thr Asp Val - 1.0 - 5 Leu Cys Ala Val Leu Tyr Ser Asn His Asn Arg Met Gly Arg His Lys 15 10 Pro His Leu Ala Leu Lys Gln Val Glu Gln Cys Leu Lys Arg Leu Lys 30 Asn Met Asn Leu Glu Gly Ser Ile Gln Asp Leu Phe Glu Leu Phe Ser

<210> 417 <211> 161 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -108..-1

Ser Lys

<400> 417 Met Thr Ser Gly Gln Ala Arg Ala Ser Xaa Gln Ser Pro Gln Ala Leu -100 -105 Glu Asp Ser Gly Pro Val Asn Ile Ser Val Ser Ile Thr Leu Thr Leu -90 -85 -80 Asp Pro Leu Lys Pro Phe Gly Gly Tyr Ser Arg Asn Val Thr His Leu -70 Tyr Ser Thr Ile Leu Gly His Gln Ile Gly Leu Ser Gly Arg Glu Ala -55 -50 His Glu Glu Ile Asn Ile Thr Phe Thr Leu Pro Thr Ala Trp Ser Ser -35. Asp Asp Cys Ala Leu His Gly His Cys Glu Gln Val Val Phe Thr Ala -20 Cys Met Thr Leu Thr Ala Ser Pro Gly Val Phe Pro Ser Leu Tyr Ser

```
-10
His Arg Thr Val Phe Leu Thr Arg Thr Ala Thr Pro Arg Ser Gly Thr 5
10
Arg Ser Ser Gln Leu Pro Glu Met Pro Thr Gln Asn Thr Pro Lys Ile
25
Thr Ile Leu Ser Gly Val Ile Arg Gly Pro Leu Glu Lys Ser Ile Met
40
Leu
```

<210> 418 <211> 67 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 418 Met Leu Gly Gly Asp His Arg Ala Leu Leu Leu Lys Ile Trp Leu Leu -15 -10 Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro Gly Arg. Leu Val Val 1 Met Glu Arg Arg Val Lys Asn Asp Leu Met Ser Phe Leu Ser Thr Val 20 25 1.5 Leu Leu Ser Phe His Ser Ser Asn Ala Arg Val Ser His Cys Glu Pro 3.0 Leu Arg Met 45

<210> 419
<211> 332
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -32..-1

<400> 419 Met Ile Xaa Leu Arg Asp Thr Ala Ala Ser Leu Arg Leu Glu Arg Asp -25 Thr Arg Gln Leu Pro Leu Leu Thr Ser Ala Leu His Gly Leu Gln Gln -10 Gln His Pro Ala Phe Ser Gly Val Ala Arg Leu Ala Lys Arg Trp Val 10 Arg Ala Gln Leu Leu Gly Glu Gly Phe Ala Asp Glu Ser Leu Asp Leu 25 Val Ala Ala Ala Leu Phe Leu His Pro Glu Pro Phe Thr Pro Pro Ser 40 Ser Pro Gln Val Gly Phe Leu Arg Phe Leu Phe Leu Val Ser Thr Phe Asp Trp Lys Asn Asn Pro Leu Phe Val Asn Leu Asn Asn Glu Leu Thr 70 75 Val Glu Glu Gln Val Glu Ile Arg Ser Gly Phe Leu Ala Ala Arg Ala 85: ..... 90 Gln Leu Pro Val Met Val Ile Val Thr Pro Gln Xaa Arg Lys Asn Ser 100

Val Trp Thr Gln Asp Gly Pro Ser Ala Gln Ile Leu Gln Gln Leu Val 120 125 Val Leu Ala Ala Glu Xaa Leu Pro Met Leu Xaa Xaa Gln Leu Met Asp 135 140 Pro Arg Gly Pro Gly Asp Ile Arg Thr Xaa Phe Arg Pro Pro Leu Asp 150 155 Ile Tyr Asp Val Leu Ile Arg Leu Ser Pro Arg His Ile Pro Arg His 165 . 175 170 Arg Gln Ala Val Asp Ser Pro Ala Ala Ser Phe Cys Arg Gly Leu Leu 180 185 190 Ser Gln Pro Gly Pro Ser Ser Leu Met Pro Val Leu Gly Xaa Asp Pro 205 200 Pro Gln Leu Tyr Leu Thr Gln Leu Xaa Glu Ala Phe Gly Asp Leu Ala 220 215 Leu Phe Phe Tyr Asp Gln His Gly Glu Val Ile Gly Val Leu Trp 230 235 Lys Pro Thr Ser Phe Gln Pro Gln Pro Phe Lys Ala Ser Ser Thr Lys 250 245 Gly Arg Met Val Met Ser Arg Gly Glu Leu Val Met Val Pro Asn 265 260 Val Glu Ala Ile Leu Glu Asp Phe Ala Val Leu Gly Glu Gly Leu Val 280 285 Gln Thr Val Glu Ala Arg Ser Glu Arg Trp Thr Val 295

<210> 420 <211> 65 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1

<210> 421 <211> 57 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1

-10 -5 1
Arg Val Tyr His Tyr Phe Gln Trp Arg Arg Ala Gln Arg Gln Ala Ala
5 10 15
Glu Glu Gln Lys Xaa Ser Gly Ile Met
20 25

<210> 422 <211> 85 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -17..-1 <400> 422 Met Lys Lys Val Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val -15 -10 Gly Phe Pro Val Ser Gln Asp Gln Glu Arg Glu Lys Arg Ser Ile Ser 10 Asp Ser Asp Glu Leu Ala Ser Gly Xaa Phe Val Phe Pro Tyr Pro Tyr 20 25 Pro Phe Arg Pro Leu Pro Pro Ile Pro Phe Pro Arg Phe Pro Trp Phe 40 35 Arg Arg Asn Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Thr Thr Pro

<210> 423 <211> 85 <212> PRT <213> Homo sapiens

55

<220>
<221> SIGNAL
<222> -17..-1

50 Leu Pro Ser Glu Lys

65

. . . . . . . . . .

<210> 424 <211> 69 <212> PRT <213> Homo sapiens

```
<220>
<221> SIGNAL
<222> -29..-1
<400> 424
Met Thr Cys Arg Gly Ser Cys Ser Tyr Ala Thr Arg Arg Ser Pro Ser
                                 -20 -15
Glu Leu Ser Leu Leu Pro Ser Ser Leu Trp Val Leu Ala Thr Ser Ser
                      -5
          -10
Pro Thr Ile Thr Ile Ala Leu Ala Met Ala Ala Gly Asn Leu Cys Pro
                                      15
                   10
Leu Pro Ser Ser Xaa Arg Xaa Lys Arg Arg Trp Cys Gln Ala Xaa Gln
               25
Gln Xaa Ala Leu Leu
              40
<210> 425
<211> 122
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -56..-1
<400> 425
Met Val Pro Trp Pro Arg Gly Lys Val Lys Thr Ala Pro Ile Pro Ile
                  -50
Ser Arg Phe Pro Phe Leu Pro Thr His Asp Pro Pro Thr Pro Ala His
                                   -30
               -35
Trp Ser Pro Ala Ser His Gln Gln Phe Lys His Xaa Ser Pro Leu Leu
                                           -10
                                  -15
               -20
Thr Leu Ala Leu Leu Gly Gln Cys Ser Leu Phe Xaa Asn Leu Arg Lys
                              1
           - 5
Lys Leu Ala Gly Gln Lys Ala Lys Lys Leu Pro Ser Phe Ser Ser Leu
                                        20
                      15
Pro Leu Thr Leu Trp Pro Leu Thr Pro Gln Phe Ala Glu Leu Thr Thr
                                     35
                  30
Val Ala Gln Lys Lys Leu Arg Trp Ser Gly Thr Leu Gly Trp Gly Pro
              4.5
Val Pro Ser Trp Val Gln Phe Phe Leu Gly
          60
<210> 426
<211> 41
 <212> PRT
<213> Homo sapiens
<220>
 <221> SIGNAL
 <222> -30..-1
```

 Arg Cys Ser Gly Ser Pro Leu Pro Leu 5 10

<210 > 427 <211 > 50 <212 > PRT <213 > Homo sapiens <220 > <221 > SIGNAL <222 > -36..-1

<210> 428 <211> 136 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1

<400> 428 Met Asp Ser Leu Arg Lys Met Leu Ile Ser Val Ala Met Leu Gly Ala -10 -15 Xaa Ala Gly Val Gly Tyr Ala Leu Leu Val Ile Val Thr Pro Gly Glu 10 Arg Arg Lys Gln Glu Met Leu Lys Glu Met Pro Leu Gln Asp Pro Arg 20 25 Ser Arg Glu Glu Ala Ala Arg Thr Gln Gln Leu Leu Leu Ala Thr Leu 40 Gln Glu Ala Ala Thr Thr Gln Glu Asn Val Ala Trp Arg Lys Asn Trp 55 Met Val Gly Gly Gly Gly Ala Thr Gly Xaa His Arg Glu Thr Gly 70 Leu Ala Ser Val Gly Ala Gly Pro Trp Leu Gly Arg Arg Asn Pro Arg 90 85 Gln Leu Ser Pro Ser Trp Ala Xaa Arg Lys Ile Arg Xaa Glu Asn Xaa 105 100 Met Pro Gly Leu Ser Gly Val Leu 115

<210> 429 <211> 194 <212> PRT <213> Homo sapiens

<220>

<221> SIGNAL <222> -65..-1

<400> 429 Met Gln Asp Ala Pro Leu Ser Cys Leu Ser Pro Thr Lys Trp Ser Ser -55 -60 Val Ser Ser Ala Asp Ser Thr Glu Lys Ser Ala Ser Ala Ala Gly Thr -35 -40 -45 Arg Asn Leu Pro Phe Gln Phe Cys Leu Arg Gln Ala Leu Arg Met Lys -25 -30 Ala Ala Gly Ile Leu Thr Leu Ile Gly Cys Leu Val Thr Gly Val Glu -10 -5 -15 Ser Lys Ile Tyr Thr Arg Cys Lys Leu Ala Lys Ile Phe Ser Arg Ala 10 5 Gly Leu Asp Asn Xaa Arg Gly Phe Ser Leu Gly Asn Trp Ile Cys Met 25 20 Ala Tyr Tyr Glu Ser Gly Tyr Asn Thr Thr Ala Gln Thr Val Leu Asp 40 Asp Gly Ser Ile Asp Tyr Gly Ile Phe Gln Ile Asn Ser Phe Ala Trp Cys Arg Arg Gly Lys Leu Lys Glu Asn Asn His Cys His Val Ala Cys 70 Ser Ala Leu Xaa Thr Asp Asp Leu Thr Asp Ala Ile Ile Cys Ala Xaa 90 85 Lys Ile Val Lys Glu Thr Gln Gly Met Asn Tyr Trp Gln Gly Trp Lys 105 100 Lys His Cys Glu Gly Arg Asp Leu Ser Xaa Trp Lys Lys Gly Cys Glu 120 Val Ser

<210> 430 <211> 141 <212> PRT <213> Homo sapiens

<220>
<221> SIGNAL
<222> -69..-1

<400> 430 Met Thr Ser Gln Pro Val Pro Asn Glu Thr Ile Ile Val Leu Pro Ser -60 -65 Asn Val Ile Asn Phe Ser Gln Ala Glu Lys Pro Glu Pro Thr Asn Gln -45 Gly Gln Asp Ser Leu Lys Lys His Leu His Ala Glu Ile Lys Val Ile -25 -30 Gly Thr Ile Gln Ile Leu Cys Gly Met Met Val Leu Ser Leu Gly Ile -10 - 1.5 Ile Leu Ala Ser Ala Ser Phe Ser Pro Asn Phe Thr Gln Val Thr Ser 5 Thr Leu Leu Asn Ser Ala Tyr Pro Phe Ile Gly Pro Phe Phe Val Xaa 20 Lys Xaa Ser Glu Glu Gly Arg Met Gly Gln Xaa Gly Glu Glu Xaa Xaa 35 Asn Ser Leu Asn Phe Pro Xaa Ala Ser Leu Leu Xaa Leu Ile Cys Gln 50 Xaa Gln Gly Phe Asn Gly Glu Ser Cys Ser Pro Val Gly

```
<210> 431
<211> 248
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -69..-1
<400> 431
Met Thr Ser Gln Pro Val Pro Asn Glu Thr Ile Ile Val Leu Pro Ser
                                 -60
Asn Val Ile Asn Phe Ser Gln Ala Glu Lys Pro Glu Pro Thr Asn Gln
                             -45
Gly Gln Asp Ser Leu Lys Lys His Leu His Ala Glu Xaa Lys Val Ile
                                            -25
               -30
Gly Thr Ile Gln Ile Leu Cys Gly Met Met Val Leu Ser Leu Gly Ile
                    -15 -10
Ile Leu Ala Ser Ala Ser Phe Ser Pro Asn Phe Thr Gln Val Thr Ser
                      5
                 1
Thr Leu Leu Asn Ser Ala Tyr Pro Phe Ile Gly Pro Phe Phe Phe Ile
        1.5
                          2.0
Ile Ser Gly Ser Leu Ser Ile Ala Thr Lys Lys Arg Leu Thr Asn Leu
      3.0
                          3.5
Leu Val His Thr Thr Leu Val Gly Ser Ile Leu Ser Ala Leu Ser Ala
                      50
Leu Val Gly Phe Ile Xaa Leu Ser Val Lys Gln Ala Thr Leu Asn Pro
                                     70 -
                  65
Ala Ser Leu Xaa Cys Glu Leu Xaa Lys Asn Asn Ile Pro Thr Xaa Xaa
                                 85
Tyr Val Xaa Tyr Phe Tyr His Asp Ser Leu Tyr Thr Asp Xaa Tyr
                             100
                                                105
Thr Ala Lys Ala Xaa Leu Ala Gly Thr Leu Ser Leu Met Leu Ile Cys
                                            120
       110
                         115
Thr Leu Leu Glu Phe Cys Xaa Xaa Val Leu Thr Ala Val Leu Arg Trp
                                        135
                     130
Lys Gln Ala Tyr Ser Asp Phe Pro Gly Ser Val Leu Phe Leu Pro Xaa
                  145 150
Ser Tyr Ile Gly Asn Ser Gly Met Ser Ser Lys Met Thr His Asp Cys
                                165
              160
Gly Tyr Glu Glu Leu Leu Thr Ser
          175
```

-25

Cys Leu Leu Ser Phe Gln Val Phe Lys Lys Lys Arg Lys Leu Xaa Leu 5

-10

```
<210> 432
<211> 49
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -36..-1
<400> 432
Met Gln Val Pro His Leu Arg Val Trp Thr Gln Val Xaa Asp Thr Phe
                   -30
Ile Gly Tyr Arg Asn Leu Gly Phe Thr Ser Met Cys Ile Leu Phe His
                 -15
```

Phe

```
<210> 433
  <211> 86
  <212> PRT
  <213> Homo sapiens
  <220>
  <221> SIGNAL
  <222> -14..-1
  <400> 433
 Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp Cys
               -10
                                     -5
 Asp Pro Arg Arg Ile His Ser Gln Asp Asp Val Leu Arg Ser Ser Ala
                             10
 Ala Asp Thr Gly Ser Ala Met Gln Arg Arg Glu Ala Trp Ala Gly Trp
                        25
. Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg Leu
                     40
                                         4.5
 Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu Gly
 His Arg Ile Cys Asp Leu
```

-318-

<210> 434 <211> 144 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -58..-1

70

<400> 434 Met Thr Arg Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro Ile -50 Pro Val Pro Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser Pro -35 Val Arg Pro Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu Leu -20 -15 Asp Ser Val Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala Val Phe Ser Thr Thr Gly Pro Ala Leu Leu Leu Leu Leu Val Ser Phe Leu Thr Phe Asp Leu Leu His Arg Pro Ala Val Thr Leu Cys His Ser Ala 30 Asn Phe Ser Pro Gly Ala Arg Val Arg Gly Pro Val Lys Val Leu Asp 45 Ser Arg Arg Leu Tyr Ser Cys Lys Trp Val Gln Ser Gln Asp Asn Leu 60 65 Ala Ser Arg Lys His Cys Cys Cys Cys Ser Trp Gly Trp Ala Arg Ser

<212> PRT <213> Homo sapiens

<220>

<221> SIGNAL <222> -16..-1

<400> 435

 Met Glu Arg Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala -15
 -10
 -5

 Ser Ala Gly Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln 1
 5
 10
 15

 Cys Phe Lys Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser 20
 25
 30

 Pro Leu Asp Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Ser 35
 40
 45

 Glu Ser Pro Pro Gly Arg Gly Xaa Val Pro Xaa Ala Gly Glu Xaa Pro

50 55 60
Val Pro Pro Pro Leu Xaa Asp Leu Xaa Met Thr Pro Arg Xaa Xaa Arg

65 70 75 80

Ala Trp Gly Pro Val Gly Pro Lys Val Pro Pro Ala Val Ser Pro Ala 85 90 95

Leu Gly Ser Gly Glu His Pro Xaa Xaa 100 105

<210> 436

<211> 162

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -16..-1

<400> 436

Met Glu Arg Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala
-15
-5

Ser Ala Gly Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln
1 10 15

Cys Phe Lys Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser

Pro Leu Asp Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Lys 35 40 45

Trp Ser Val Arg Val Leu Leu Ser Lys Arg Cys Ala Pro Arg Cys Pro 50 55 60

Asn Asp Asn Met Xaa Phe Glu Trp Ser Pro Ala Pro Met Val Gln Gly

70 75 80 Val Ile Thr Arg Arg Cys Cys Ser Trp Ala Leu Cys Asn Arg Ala Leu

85 90 95
Thr Pro Gln Glu Gly Arg Trp Ala Leu Xaa Gly Gly Leu Leu Leu Gln

Asp Pro Ser Arg Gly Xaa Lys Thr Trp Val Arg Pro Gln Leu Gly Leu 115 120 125

Pro Leu Cys Leu Pro Xaa Ser Asn Pro Leu Cys Pro Xaa Glu Thr Gln
130 135 140

Glu Gly

145

```
<210> 437
  <211> 110
  <212> PRT
  <213> Homo sapiens
  <220>
  <221> SIGNAL
  <222> -20..-1
  <400> 437
  Met Xaa Leu Met Val Leu Val Phe Thr Ile Gly Leu Thr Leu Leu
                  -15
                                       -10
  Gly Xaa Gln Ala Met Pro Ala Asn Arg Leu Ser Cys Tyr Arg Lys Ile
  Leu Lys Asp His Asn Cys His Asn Leu Pro Glu Gly Val Ala Asp Leu
                             20
  Thr Gln Ile Asp Val Asn Val Gln Asp His Phe Trp Asp Gly Lys Gly
                                            40
  Cys Glu Met Ile Cys Tyr Cys Asn Phe Lys Arg Ile Ala Leu Leu Pro
                    50
                                        55
. Lys Arg Arg Phe Leu Trp Thr Lys Asp Leu Phe Arg Asp Ser Leu Gln
                                     70
  Gln Ser Met Arg Ile Phe Met Tyr Ser Gly Glu His His Ser
                                85
 <210> 438
 <211> 71
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -15..-1
 <400> 438
 Met Lys Leu Leu Thr His Asn Leu Leu Ser Ser His Val Arg Gly Val
               -10
                              -5
 Gly Ser Arg Gly Phe Pro Leu Arg Leu Gln Ala Thr Glu Val Arg Ile
                                10
 Cys Pro Val Glu Phe Asn Pro Asn Phe Val Ala Arg Met Ile Pro Lys
                           25
 Val Glu Trp Ser Ala Phe Leu Glu Ala Xaa Asp Asn Leu Arg Leu Ile
 Gln Val Pro Arg Arg Ala Gly
 <210> 439
 <211> 99
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -24..-1
 Met Lys Ser Ala Lys Leu Gly Phe Leu Leu Arg Phe Phe Ile Phe Cys
```

-20

 Ser
 Leu
 Asn
 Thr
 Leu
 Leu
 Leu
 Gly
 Gly
 Val
 Asn
 Lys
 Ile
 Ala
 Glu
 Lys

 Ile
 Cys
 Gly
 Asp
 Leu
 Lys
 Asp
 Pro
 Cys
 Lys
 Leu
 Asp
 Met
 Asn
 Phe
 Gly

 Ser
 Cys
 Tyr
 Glu
 Val
 His
 Phe
 Arg
 Tyr
 Phe
 Tyr
 Asn
 Arg
 Thr
 Ser
 Lys

 Arg
 Cys
 Glu
 Thr
 Phe
 Val
 Phe
 Ser
 Ser
 Cys
 Asn
 Gly
 Asn
 Leu
 Asn
 Asn
 Asn

 Phe
 Lys
 Leu
 Lys
 Ile
 Glu
 Arg
 Glu
 Val
 Xaa
 Cys
 Val
 Ala
 Lys
 Tyr
 Lys

 Phe
 Lys
 Leu
 Arg
 Glu
 Val
 Xaa
 Cys
 Val
 Ala
 Lys
 Tyr
 Lys

 Pro
 Pro
 Arg
 France
 France
 France

<210> 440 <211> 169 <212> PRT <213> Homo sapiens . <220> <221> SIGNAL <222> -25..-1 <400> 440 Met Arg Lys Pro Ala Ala Gly Phe Leu Pro Ser Leu Leu Lys Val Leu -20 -15 Leu Leu Pro Leu Ala Pro Ala Ala Ala Gln Asp Ser Thr Gln Ala Ser - 5 Thr Pro Gly Ser Pro Leu Ser Pro Thr Glu Tyr Gln Arg Phe Phe Ala 15 Leu Leu Thr Pro Thr Trp Lys Ala Glu Thr Thr Cys Arg Leu Arg Ala 3.0 35 Thr His Gly Cys Arg Asn Pro Thr Leu Val Gln Leu Asp Gln Tyr Glu 45 50 Asn His Gly Leu Val Pro Asp Gly Ala Val Cys Ser Asn Leu Pro Tyr 60 65 Ala Ser Trp Phe Glu Ser Phe Cys Gln Phe Thr His Tyr Arg Cys Ser 80 Asn His Val Tyr Tyr Ala Lys Arg Val Leu Cys Ser Gln Pro Val Ser 95 Ile Leu Ser Pro Asn Thr Leu Lys Glu Ile Glu Xaa Ser Ala Glu Val 110 115 Ser Pro Thr Thr Asp Asp Leu Pro His Leu Thr Pro Leu His Ser Asp 125 130 Arg Thr Pro Asp Leu Pro Ala Leu Ala 140

<211> 167
<212> PRT
<213> Homo sapiens

<220>
<221> SIGNAL
<222> -76..-1

<400> 441

Met Gly Asp Tyr Leu Leu Arg Gly Tyr Arg Met Leu Gly Glu Thr Cys
-75
-70
-65

<210> 441

Ala Asp Cys Gly Thr Ile Leu Leu Gln Asp Lys Gln Arg Lys Ile Tyr -50 -55 Cys Val Ala Cys Gln Glu Leu Asp Ser Asp Val Asp Lys Asp Asn Pro -35 -40 Ala Leu Asn Ala Gln Ala Ala Leu Ser Gln Ala Arg Glu His Gln Leu -25 -20 Ala Ser Ala Ser Glu Leu Pro Leu Gly Ser Arg Pro Ala Pro Gln Pro -10 -5 Pro Val Pro Arg Pro Glu His Cys Glu Gly Ala Ala Ala Gly Leu Lys 10 15 Ala Ala Gln Gly Pro Pro Ala Pro Ala Val Pro Pro Asn Thr Xaa Val 30 Met Ala Cys Thr Gln Thr Ala Leu Leu Gln Lys Leu Thr Trp Ala Ser 45 Ala Glu Leu Gly Ser Xaa Thr Ser Xaa Gly Lys Xaa Ala Ser Ser Cys 60 Val Ala Leu Ser Ala His Val Arg Arg Pro Cys Ala Ala Cys Ser Ser 75 Tyr Ser Thr Lys Arg Ser Pro 85 9.0

<210> 442 <211> 70 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 442 Met Ile Leu Cys Phe Leu Leu Pro His His Arg Leu Gln Glu Ala Arg -10 - 5 Gln Ile Gln Val Leu Lys Met Leu Pro Arg Glu Lys Leu Arg Arg Arg 10 Glu Glu Arg Lys Gln Ile Asn Gly Lys Lys Xaa Arg Thr Lys Tyr Glu 25 30 Thr Pro Arg Lys Xaa Xaa Gly Lys Lys Gly Gly Asn Xaa Xaa Xaa Xaa Xaa Leu Ser Lys Arg Asp

<210> 443 <211> 381 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -33..-1 <400> 443 Met Ser Trp Thr Val Pro Val Val Arg Ala Ser Gln Arg Val Ser Ser -30 -25 -20 Val Gly Ala Asn Xaa Leu Cys Leu Gly Met Ala Leu Cys Pro Arg Gln -10 - 5 Ala Thr Arg Ile Pro Leu Asn Gly Thr Trp Leu Phe Thr Pro Val Ser

Lys Met Ala Thr Val Lys Ser Glu Leu Ile Glu Arg Phe Thr Ser Glu 20 25 Lys Pro Val His His Ser Lys Val Ser Ile Ile Gly Thr Gly Ser Val 40 Gly Met Ala Cys Ala Ile Ser Ile Leu Leu Lys Gly Leu Ser Asp Glu 55 Leu Ala Leu Val Asp Leu Asp Glu Xaa Lys Leu Lys Gly Glu Thr Met 70 Asp Leu Gln His Gly Ser Pro Phe Thr Lys Met Pro Asn Ile Val Cys 85 90 Ser Lys Xaa Tyr Phe Val Thr Ala Asn Ser Asn Leu Val Ile Ile Thr 100 105 Ala Gly Ala Arg Gln Xaa Lys Gly Glu Thr Arg Leu Asn Leu Xaa Gln 120 Arg Asn Val Ala Ile Phe Lys Leu Met Ile Ser Ser Ile Val Gln Tyr 130 135 140 Ser Pro His Cys Lys Leu Ile Ile Val Ser Asn Pro Val Asp Ile Leu 150 Thr Tyr Val Ala Trp Lys Leu Ser Ala Phe Pro Lys Asn Arg Ile Ile 165 170 Gly Ser Gly Cys Asn Leu Ile Xaa Ala Arg Phe Arg Phe Leu Ile Gly 180 185 Gln Lys Leu Gly Ile His Ser Glu Sar Cys His Gly Trp Ile Leu Gly 200 Glu His Gly Asp Ser Ser Val Pro Val Trp Ser Gly Val Asn Ile Ala 215 Gly Val Pro Leu Lys Asp Leu Asn Ser Asp Ile Gly Thr Asp Lys Asp 230 Pro Glu Gln Trp Lys Asn Val His Lys Glu Val Thr Ala Thr Ala Tyr 245 Glu Ile Ile Lys Met Lys Gly Tyr Thr Ser Trp Ala Ile Gly Leu Ser 260 265 Val Ala Asp Leu Thr Glu Ser Ile Leu Lys Asn Leu Arg Arg Ile His 280 Pro Val Ser Thr Ile Thr Lys Gly Leu Tyr Gly Ile Xaa Glu Glu Val 295 Phe Leu Ser Ile Pro Cys Ile Leu Gly Glu Asn Gly Ile Thr Asn Leu 310 Ile Lys Ile Lys Leu Thr Pro Glu Glu Glu Ala His Leu Lys Lys Ser 325 330 Ala Lys Thr Leu Trp Glu Ile Gln Asn Lys Leu Lys Leu

```
<210> 445
<211> 50
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -37..-1
<400> 445
Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val Asn
     -35
                    -30
Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala Leu
                                       -10
                      -15
Ser Pro Cys Leu Thr Ala Pro Lys Ser Pro Arg Leu Ala Met Met Pro
                  1
Asp Asn
<210> 446
<211> 51
<212> PRT
<213 > Homo sapiens
<220>
<221> SIGNAL
<222> -26..-1
<400> 446
Met Thr Pro Trp Cys Leu Ala Cys Leu Gly Arg Arg Pro Leu Ala Ser
                             -15
                -20
Leu Gln Trp Ser Leu Thr Leu Ala Trp Cys Gly Ser Gly Ser His Trp
                  -5
                                      1
Thr Glu Arg Pro Xaa Gln Xaa Ser Pro Trp Xaa Ser Leu Ser Ala Thr
Thr Arg Gly
       25
<210> 447
<211> 242
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -30..-1
<400> 447
Met Gly Glu Ala Ser Pro Pro Ala Pro Ala Arg Arg His Leu Leu Val
                  -25
                                      -20
Leu Leu Leu Leu Ser Thr Leu Val Ile Pro Ser Ala Ala Ala Pro
               -10
                                  -5
Ile His Asp Ala Asp Ala Gln Glu Ser Ser Leu Gly Leu Thr Gly Leu
                          10
Gln Ser Leu Leu Gln Gly Phe Ser Arg Leu Phe Leu Lys Gly Asn Leu
                       25
Leu Arg Gly Ile Asp Ser Leu Phe Ser Ala Pro Met Asp Phe Arg Gly
```

-50

15

30

-20

Gln Ser Cys Ile Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met 45 Cys Asn Ala Leu Cys Leu Phe Gln Val Pro Lys Ser Ser Leu Ala Phe

Gln Thr Asp Gly Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln

75

40 45 Leu Pro Gly Asn Tyr His Lys Glu Glu Asn Gln Glu His Gln Leu Gly 55 60 Asn Asn Thr Leu Ser Ser His Leu Gln Ile Asp Lys Met Thr Asp Asn 70 75 Lys Thr Gly Glu Val Leu Ile Ser Glu Asn Val Val Ala Ser Ile Gln 90 Pro Xaa Glu Gly Xaa Phe Glu Gly Asp Leu Lys Val Pro Arg Met Glu 105 110 Glu Lys Glu Ala Leu Val Pro Xaa Gln Lys Ala Thr Asp Ser Phe His 120 125 Thr Glu Leu His Pro Arg Val Ala Phe Trp Ile Ile Lys Leu Pro Arg 135 140 Arg Arg Ser His Gln Asp Ala Leu Glu Gly Gly His Trp Leu Xaa Glu 155 Lys Arg His Arg Leu Gln Ala Ile Arg Asp Gly Leu Arg Lys Gly Thr 170 175 His Lys Asp Xaa Leu Xaa Xaa Gly Thr Glu Ser Ser His Ser Arg 185 190 Leu Ser Pro Arg Lys Xaa His Leu Leu Tyr Ile Leu Xaa Pro Ser Arg 195 200 205 Gln Leu

<210> 448 <211> 154 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -60..-1 <400> 448 Met Gly Ser Lys Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu -55 Arg Gln Arg Arg Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys -40 -35 Arg Val Lys Ala Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu Val Arg Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln <del>-</del>5 · Cys Trp Trp Arg Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln 10 Ala Leu Leu Gly Val Tyr Val Ile Gln Glu Gln Ala Ala Val Lys Leu

90

Pro Glu Phe His Ile Glu Ile Leu Ser Ile

<210> 449 <211> 89 <212> PRT <213> Homo sapiens

```
<220>
<221> SIGNAL
<222> -61..-1
<400> 449
Met Asn Ala Ala Ile Asn Thr Gly Pro Ala Pro Ala Val Thr Lys Thr
               -55
                            -50
Glu Thr Glu Val Gln Asn Pro Asp Val Leu Trp Asp Leu Asp Ile Pro
                 -40
                                  -35
Glu Ala Arg Ser His Ala Asp Gln Asp Ser Asn Pro Lys Ala Glu Ala
             -25
                              -20
Leu Leu Pro Cys Asn Leu His Cys Ser Trp Leu His Ser Ser Pro Arg
          -10 -5
Pro Asp Pro His Ser His Phe Pro Ser Xaa Arg Arg Cys Pro Leu Pro
           10
His Pro Cys Ala Thr Tyr Pro Pro Xaa
20 25
<210> 450
<211> 73
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -26..-1
<400> 450
Met Arg Met Ser Leu Ala Gln Arg Val Leu Leu Thr Trp Leu Phe Thr
 -25
                    -20
                                     -15
Leu Leu Phe Leu Ile Met Leu Val Leu Lys Leu Asp Glu Lys Ala Pro
-10 -5
Trp Asn Trp Phe Leu Ile Phe Ile Pro Val Trp Ile Phe Asp Thr Ile
   10
                        15
                                   20
Leu Leu Val Leu Leu Ile Val Lys Met Ala Gly Arg Cys Lys Ser Gly
 25 30
                                   35
Phe Asp Leu Asp Met Asp His Thr Ile
                    45
<210> 451
<211> 54
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -34..-1
<400> 451
Met Ile Pro Leu Ile Ser His Leu Ala Glu Ala Ala Pro Pro Thr Ser
                              -25 -20
Trp Ser Leu Ile Ser Ser Val Leu Asn Val Gly His Leu Leu Phe Ser
                          -10
                                     -5
Ser Ala Cys Ser Val Ser Leu Glu Ala Leu Ser Thr Arg Asn Ile Lys
    1 5
Ala Ile Ile Leu Met Lys
            20
```

```
<210> 452
<211> 121
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -38..-1
<400> 452
Met Glu Ser Pro Gln Leu His Cys Ile Leu Asn Ser Asn Ser Val Ala
                              -30
Cys Ser Phe Ala Val Gly Ala Gly Phe Leu Ala Phe Leu Ser Cys Leu
                          -15
                                             -10
Ala Phe Leu Val Leu Asp Thr Gln Glu Thr Arg Ile Ala Gly Thr Arg
Phe Lys Thr Ala Phe Gln Leu Leu Asp Phe Ile Leu Ala Val Leu Trp
         15
                                 20
Ala Val Val Trp Phe Met Gly Phe Cys Phe Leu Ala Asn Gln Trp Gln
        3.0
                           35
His Ser Pro Pro Lys Glu Xaa Leu Leu Gly Ser Ser Ser Ala Gln Ala
    4.5
                   50
Ala Ile Gly Xaa His Leu Leu His Pro Cys Leu Asp Ile Pro Xaa .
                    6.5
Leu Pro Gly Xaa Pro Gly Pro Pro Lys
<210> 453
<211> 166
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -37..-1
<400> 453
Met Ser Thr Val Gly Leu Phe His Phe Pro Thr Pro Leu Thr Arg Ile
                          -30
Cys Pro Ala Pro Trp Gly Leu Arg Leu Trp Glu Lys Leu Thr Leu Leu
                      -15
                                      -10
Ser Pro Gly Ile Ala Val Thr Pro Val Gln Met Ala Gly Lys Lys Asp
                  1
                               5
Tyr Pro Ala Leu Leu Ser Leu Asp Glu Asn Glu Leu Glu Glu Gln Phe
           15 20
Val Lys Gly His Gly Pro Gly Gly Gln Ala Thr Asn Lys Thr Ser Asn
                         35
                                             40
Cys Val Val Leu Lys Xaa Ile Pro Ser Gly Ile Val Val Lys Cys His
Gln Thr Arg Ser Val Asp Gln Asn Arg Lys Leu Ala Arg Lys Ile Leu
                                     70
Gln Glu Lys Val Xaa Val Phe Tyr Asn Gly Glu Asn Ser Pro Val His
                                 85
Lys Glu Lys Arg Glu Ala Ala Lys Lys Gln Glu Arg Lys Lys Arg
          95
                             100
Ala Lys Glu Thr Leu Glu Lys Lys Xaa Leu Lys Xaa Leu Trp Glu
                         115
```

Ser Ser Lys Lys Val His 125

<210> 454 <211> 180 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 454 Met Gly Ile Gln Thr Ser Pro Val Leu Leu Ala Ser Leu Gly Val Gly ~20 Leu Val Thr Leu Leu Gly Leu Ala Val Gly Ser Tyr Leu Val Arg Arg - 5 Ser Arg Arg Pro Gln Val Thr Leu Leu Asp Pro Asn Glu Lys Tyr Leu 15 Leu Arg Leu Leu Asp Lys Thr Thr Val Ser His Asn Thr Lys Arg Phe 30 Arg Phe Ala Leu Pro Thr Ala His His Thr Leu Gly Leu Pro Val Gly 45 Lys His Ile Tyr Leu Ser Thr Arg Ile Asp Gly Ser Leu Val Ile Arg 60 65 Pro Tyr Thr Pro Val Thr Ser Asp Glu Asp Gln Gly Tyr Val Asp Leu 75 8.0 Val Xaa Lys Val Tyr Leu Lys Gly Val His Pro Lys Phe Pro Glu Gly 95 Gly Lys Met Ser Xaa Tyr Leu Asp Xaa Leu Lys Val Gly Asp Xaa Val 110 115 Glu Phe Xaa Gly Pro Ser Gly Leu Leu Thr Tyr Thr Gly Lys Gly His 125 130 Phe Asn Ile Gln Pro Asn Lys Asn Leu His Gln Asn Pro Glu Trp Arg 145 Arg Asn Trp Glu

<210> 455
<211> 91
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -64..-1

Phe Pro Arg Phe Tyr Phe Leu Cys Leu Ser Ser Leu Thr Ala Phe Gly 1 5 10 15

Thr Pro Pro Ile Gly Val His Ile Pro Ser Pro

25

20

```
<210> 456
<211> 257
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -23..-1
<400> 456
Met Arg Arg Ile Ser Leu Thr Ser Ser Pro Val Arg Leu Leu Leu Xaa
    -20 -15 -10
Leu Leu Leu Leu Ile Ala Leu Glu Ile Met Val Gly Gly His Ser
    - 5
                       1
Leu Cys Phe Asn Phe Thr Ile Lys Ser Leu Ser Arg Pro Gly Gln Pro
10
     15
Trp Cys Glu Ala His Val Phe Leu Asn Lys Asn Leu Phe Leu Gln Tyr
           30
                     35
Asn Ser Asp Asn Asn Met Val Lys Pro Leu Gly Leu Leu Gly Lys Lys
       4.5
              50
Val Tyr Ala Thr Ser Thr Trp Gly Glu Leu Thr Gln Thr Leu Gly Glu
                       65
Val Gly Arg Asp Leu Arg Met Leu Leu Cys Asp Ile Lys Pro Gln Ile
             80
Lys Thr Ser Asp Pro Ser Thr Leu Gln Val Xaa Xaa Phe Cys Gln Arg
90
          95
                                 100
Glu Ala Glu Arg Cys Thr Gly Ala Ser Trp Gln Phe Ala Thr Asn Gly
             110
                              115
Glu Lys Ser Leu Leu Phe Asp Ala Met Asn Met Thr Trp Thr Val Ile
          125
                           130
Asn His Glu Ala Ser Xaa Ile Lys Glu Thr Trp Lys Lys Asp Arg Xaa
                        145
                                        150
Leu Glu Xaa Tyr Phe Arg Lys Leu Ser Lys Gly Asp Cys Asp His Trp
                    160
Leu Arg Glu Phe Leu Gly His Trp Glu Ala Met Pro Xaa Pro Xaa Val
                175
                               180 185
Ser Pro Xaa Asn Ala Ser Xaa Ile His Trp Ser Ser Ser Xaa Leu Pro
                               195
Xaa Xaa Trp Ile Ile Leu Gly Ala Phe Ile Leu Leu Xaa Leu Met Gly
        205 210
Ile Val Leu Ile Cys Val Trp Trp Gln Asn Gly Xaa Xaa Ser Thr Xaa
                       225
Xaa
```

```
<210> 457
<211> 193
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -60..-1
```

<400> 457
Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu Pro
-60 -55 -50 -45

Cys Ser Gly Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile Pro -40 -35 Leu Leu Cly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val Leu -25 -20 Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp Pro - 5 Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala Pro 10 Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg Ala 30 Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Pro Xaa Thr 45 Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile Val 60 Xaa Cys Leu His Leu His Xaa Leu Leu Ser Val Glu Thr Arg Xaa Phe 75 Xaa Lys His Leu Leu Val Leu Val Ala Val Ala His Ser Val Leu 90 95 Glu Pro Pro Ala Leu Val Pro Asn Val Gln Cys Glu Met Cys Thr His 110 Ser Gly Pro Arg Asp Leu Glu Ala Ala Val Val Ser Pro Ala Pro Trp Glu

<210> 458 <211> 107 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 458 Met Val Leu Thr Leu Gly Glu Ser Trp Pro Val Leu Val Gly Arg Arg -25 -20 -15 Phe Leu Ser Leu Ser Ala Ala Asp Gly Ser Asp Gly Ser His Asp Ser -10 Trp Asp Val Glu Arg Val Ala Glu Trp Pro Trp Leu Ser Gly Thr Ile 10 15 Arg Ala Val Ser His Thr Asp Val Thr Lys Lys Asp Leu Lys Val Cys 30 Val Glu Phe Xaa Gly Glu Ser Trp Arg Lys Arg Arg Trp Ile Glu Val 45 Tyr Ser Leu Leu Arg Lys Ala Phe Leu Val Lys His Asn Leu Val Leu Ala Glu Arg Lys Ser Pro Glu Ile Ser Trp Gly

<210> 459
<211> 121
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -13..-1

<400> 459 Met Leu Val Leu Arg Ser Ala Leu Thr Arg Ala Leu Ala Ser Arg Thr -5 Leu Ala Pro Gln Met Cys Ser Ser Phe Ala Thr Gly Pro Arg Gln Tyr 10 Asp Gly Ile Phe Tyr Glu Phe Arg Ser Tyr Tyr Leu Lys Pro Ser Lys 25 30 Met Asn Glu Phe Leu Glu Asn Phe Glu Lys Asn Ala Gln Leu Arg Thr 45 Ala His Ser Glu Leu Val Gly Tyr Trp Ser Val Xaa Phe Gly Gly Arg Met Xaa Thr Val Phe His Ile Trp Lys Tyr Asp Asn Phe Ala His Arg Thr Glu Phe Gln Lys Ala Leu Ala Lys Asp Lys Glu Trp Gln Glu Gln 90 Phe Leu Ile Pro Asn Leu Ala Leu Asn 100 . 105

<210> 461 <211> 109 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -13..-1

85

90

95

```
<210> 462
<211> 143
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -41..-1
<400> 462
Met Ala Thr Ala Thr Glu Gln Trp Val Leu Val Glu Met Val Gln Ala
                        -35
                                           -30
Leu Tyr Glu Ala Pro Ala Tyr His Leu Ile Leu Glu Gly Ile Leu Ile
                   -20
                                       -15
Leu Trp Ile Ile Arg Leu Leu Phe Ser Lys Thr Tyr Lys Leu Gln Glu
                                 1
Arg Ser Asp Leu Thr Val Lys Glu Lys Glu Glu Leu Ile Glu Glu Trp
                         15
Gln Pro Glu Pro Leu Val Pro Pro Val Pro Lys Asp His Pro Ala Leu
                                         3.5
Asn Tyr Asn Ile Val Ser Gly Pro Pro Ser His Lys Thr Val Val Asn
                   45
Gly Lys Glu Cys Ile Asn Phe Ala Ser Phe Asn Phe Leu Gly Leu Leu
               60
                                   65
Asp Asn Pro Arg Val Lys Ala Ala Ala Leu Ala Ser Leu Lys Lys Tyr
           75
                        80
Gly Val Gly Thr Cys Gly Pro Cys Gly Phe Tyr Gly Thr Phe Glu
<210> 463
<211> 232
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -30..-1
<400> 463
Met Ala Ala Thr Ser Gly Thr Asp Glu Pro Val Ser Gly Glu Leu Val
                -25
                                       -20
Ser Val Ala His Ala Leu Ser Leu Pro Ala Glu Ser Tyr Gly Asn Xaa
               -10
                                   - 5
Xaa Asp Ile Glu Met Ala Trp Ala Met Arg Ala Met Gln His Ala Glu
                           10
Val Tyr Tyr Lys Leu Ile Ser Ser Val Asp Pro Gln Phe Leu Lys Leu
Thr Lys Val Asp Asp Gln Ile Tyr Ser Glu Phe Arg Lys Asn Phe Glu
                   40
Thr Leu Arg Ile Asp Val Leu Xaa Pro Glu Xaa Leu Lys Ser Glu Ser
```

60

Ala Lys Glu Pro Pro Gly Tyr Asn Ser Leu Pro Leu Lys Leu Leu Gly
70 75 80

Thr Gly Lys Ala Ile Thr Lys Leu Phe Ile Ser Val Phe Arg Thr Lys
85 90 95

Lys Glu Arg Lys Glu Ser Thr Met Glu Glu Lys Lys Glu Leu Thr Val

<220>

```
105
   100
Glu Lys Lys Arg Thr Pro Arg Met Glu Glu Arg Lys Glu Leu Ile Val
           120
                                  125
Glu Lys Lys Lys Arg Lys Glu Ser Thr Glu Lys Thr Lys Leu Thr Lys
              135
                               140
                                              145
Glu Glu Lys Lys Gly Lys Lys Leu Thr Lys Lys Ser Thr Lys Val Val
                         155
                                             160
Lys Lys Leu Cys Lys Val Tyr Arg Glu Gln His Ser Arg Ser Tyr Asp
                      170
                                   175
Ser Ile Glu Thr Thr Ser Thr Thr Val Leu Leu Ala Gln Thr Pro Leu
              185
                             190
Val Lys Cys Lys Phe Leu Tyr Asn
       200
<210> 464
<211> 61
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 464
Met Thr Phe Arg His Gln Asp Asn Ser Leu Met Phe Phe Ser Met Met
-20 -15 -10
Ala Thr Cys Thr Ser Asn Val Gly Phe Thr His Thr Thr Met Asn Cys
Ser Leu Thr Ser Pro Val Asp Phe Lys Asp Leu Leu Arg Val Leu Leu
       15
              20
Ile Lys Phe Gly Tyr Asp Arg Lys Ser Thr Ile Lys Ser
                        35
<210> 465
<211> 34
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -19..-1
<400> 465
Met Phe Leu Lys Ser Gly Ala Gly Leu Ser Ser Cys Leu Leu Pro Leu
           -15
                             -10
Cys Trp Leu Glu Arg Lys Asp His Gly Arg Arg Pro Ser Xaa His Pro
Gly Arg
  15
<210> 466
<211> 215
<212> PRT
<213> Homo sapiens
```

<221> SIGNAL <222> -54..-1

<400> 466 Met Asn Xaa Tyr Ala Ser Pro Phe Asn Xaa Gln Leu Xaa Tyr Leu Xaa -50 Leu Ser Arg Phe Glu Cys Val His Arg Asp Gly Arg Val Ile Thr Leu -30 Ser Tyr Gln Glu Gln Glu Leu Gln Asp Phe Leu Leu Ser Gln Met Ser -15 -10 Gln His Gln Val His Ala Val Gln Gln Leu Ala Lys Val Met Gly Trp Gln Val Leu Ser Phe Ser Asn His Val Gly Leu Gly Pro Ile Glu Ser 20 Xaa Gly Asn Ala Ser Ala Ile Thr Val Ala Pro Gln Val Val Thr Met 35 Leu Phe Gln Phe Val Met Asp Leu Lys Val Ala Ala Arg Leu Trp Phe 50 Ser Phe Leu Val Thr Asn Val Lys Thr Phe Gln Lys Val Met Phe Tyr 65 Lys Ile Thr Asn Gly Val Ile Phe Val Gly His Ser Lys Lys Phe Ser 80 85 Gly Ile Lys Trp Lys Val Xaa Ile Leu Phe Ile Lys Trp Xaa Cys Leu 100 95 Cys Leu His Leu Ala Leu Val Tyr Tyr Asp Phe Phe Gln Met Phe Pro 115 110 Lys Xaa Val Ser Xaa Asn Phe Asp Leu Lys Cys Leu Gln Ile Asn Tyr 130 135 Lys His Lys Glu Glu Ile Thr Ser Lys Arg Val Leu Phe Leu Lys Ile 140 145 150 Ile Ile Arg Lys Cys Phe Ile

5

<210> 468 <211> 85 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1

<400> 468

 Met
 Cys
 Ser
 His
 Ala
 Ser
 Met
 Ser
 Phe
 His
 Thr
 Leu
 Phe
 His
 Leu
 Phe
 Leu
 Phe
 Leu
 Phe
 Lys
 Pro
 Gln
 Ser
 Lys
 His
 Cys
 Cys
 Phe
 Leu
 Phe
 Phe</th

<210> 469 <211> 51 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -16..-1 <400> 469 Met Leu Arg Ile Ala Leu Thr Leu Ile Pro Ser Met Leu Ser Arg Ala -15 -10 <del>-</del> 5 Ala Gly Trp Cys Trp Tyr Lys Glu Pro Thr Gln Gln Phe Ser Tyr Leu 10 15 Cys Leu Pro Cys Leu Ser Trp Asn Lys Lys Gly Asn Val Leu Gln Leu 25 Pro Asn Phe

<211> 67 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -43..-1 <400> 470 Met Thr Pro Gln Tyr Leu Pro His Gly Gly Lys Tyr Gln Val Leu Gly -40 -35 Asp Tyr Ser Leu Ala Val Val Phe Pro Leu His Phe Ser Asp Leu Ile -25 -20 -15 Ser Val Leu Tyr Leu Ile Pro Lys Thr Leu Thr Thr Asn Thr Ala Val -5 1

Lys His Ser Ile Gln Lys Asn Cys Met Xaa Leu Val Leu Gly Lys Leu

<210> 471 <211> 63 <212> PRT <213> Homo sapiens

Leu Ser Gln

35

<210> 470

<220>

```
<221> SIGNAL
<222> -15..-1
<400> 471
Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe Ala Arg Ala Leu
        -10
                               -5
Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser Glu Lys His Arg
                              10
Leu Glu Lys Cys Arg Glu Leu Glu Ser Ser His Ser Ala Pro Gly Ser
                          25
Thr Gln His Arg Arg Lys Thr Thr Arg Arg Asn Tyr Ser Ser Ala
                     40
<210> 472
<211> 179
<212> PRT
<213> Homo sapiens
<220>
2215 SIGNAL
<222> -58..-1
<400> 472
Met Ser Thr Gly Gln Leu Tyr Arg Met Glu Asp Ile Gly Arg Phe His
          - 5 5
                              -50
Ser Gln Gln Pro Gly Ser Leu Thr Pro Ser Ser Pro Thr Val Gly Glu
                          -35
Ile Ile Tyr Asn Asn Thr Arg Asn Thr Leu Gly Trp Ile Gly Gly Ile
                      -20
                                         -15
Leu Met Gly Ser Phe Gln Gly Thr Ile Ala Gly Gln Gly Thr Gly Ala
               <del>-</del> 5
Thr Ser Ile Ser Glu Leu Cys Lys Gly Gln Glu Leu Glu Pro Ser Gly
        10
                            15
Ala Gly Leu Thr Val Ala Pro Pro Gln Ala Val Ser Leu Gln Gly Ile
                         30
                                35
Tyr Thr Leu Pro Trp Leu Leu Gln Leu Phe His Ser Thr Ala Leu Xaa
                                        50
                     45
Xaa Xaa Gln Gln Pro Asn Gly Ser Leu Ser Leu Asn Ile Ser Ser Ser
                60
His Ala Pro Xaa Pro Xaa Thr Cys Thr Leu Glu Pro Gly Val Asp Pro
              75
                                 80
Thr Arg Xaa Val Cys Ile Asn Pro His Pro Pro Pro Pro Ile Leu Lys
                             95
Xaa Pro Leu Ser Pro Tyr Pro Lys Pro Gln Leu Gly Thr His Ala Gly
Gln Val Asn
   120
<210> 473
<211> 238
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -71..-1
```

<400> 473 Met Xaa Xaa Phe Thr Asp Pro Ser Ser Val Asn Glu Lys Lys Arg Arg -65 Glu Arg Glu Glu Arg Gln Asn Ile Val Leu Trp Arg Gln Pro Leu Ile -50 -45 Thr Leu Gln Tyr Phe Ser Leu Glu Ile Leu Val Ile Leu Lys Glu Trp -35 -30 Thr Ser Lys Leu Trp His Arg Gln Ser Ile Val Val Ser Phe Leu Leu -20 -15 Leu Leu Ala Gly Leu Ile Ala Thr Tyr Tyr Val Glu Gly Val His Gln Gln Tyr Val Gln Arg Ile Glu Lys Gln Phe Leu Leu Tyr Ala Tyr Trp 15 Ile Gly Leu Gly Ile Leu Ser Ser Val Gly Leu Gly Thr Gly Leu His 30 Thr Phe Leu Leu Tyr Leu Gly Pro His Ile Ala Ser Val Thr Leu Ala 50 Ala Tyr Glu Cys Asn Ser Val Asn Phe Pro Glu Pro Pro Tyr Pro Asp 65 Gln Ile Ile Cys Pro Asp Glu Glu Gly Thr Glu Gly Thr Ile Ser Leu 80 Trp Ser Ile Ile Ser Lys Val Arg Ile Glu Ala Cys Met Trp Gly Ile 9.5 100 Gly Thr Ala Ile Gly Glu Leu Pro Pro Tyr Phe Met Ala Arg Ala Ala 115 120 110 Arg Leu Ser Gly Ala Glu Pro Asp Asp Glu Glu Tyr Gln Glu Phe Glu 130 135 Glu Met Leu Glu His Ala Glu Ser Ala Gln Val Arg Thr Val Gly Ile 140 145 150 Glu Asn Arg Thr Leu Tyr Phe Phe Leu Lys Arg Leu Leu Arg

<210> 474 <211> 178 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1 <400> 474 Met Glu Arg Gln Ser

Met Glu Arg Gln Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe -30 -25 Gln His Gln Gly Ala Val Glu Leu Leu Val Phe Asn Phe Leu Leu Ile -10 -15 Leu Thr Ile Leu Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr Gly Gly Ala Met Val Tyr Gly Leu Xaa Met Gly Leu 20 Ile Leu Xaa Tyr Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Xaa Val 35 Tyr Asp Cys Val Lys Leu Thr Phe Ser Pro Ser Thr Leu Leu Val Asn 50 Ile Thr Asp Gln Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln 70 65 His Xaa Ile Asn Pro His Xaa Gly Asn Ala Ile Leu Glu Lys Met Thr 85 Phe Asp Pro Xaa Ile Phe Phe Asn Val Leu Leu Pro Pro Ile Ile Phe

```
105
                         100
His Ala Gly Tyr Ser Leu Lys Lys Arg His Phe Phe Gln Asn Leu Gly
110 115
                              120
Ser Ile Leu Thr Tyr Ala Phe Leu Gly Thr Ala Ile Ser Cys Ile Val
             130
                             135
Ile Gly
140
<210> 475
<211> 96
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 475
Met Ser Met Gln Phe Leu Phe Lys Met Val Ala Leu Cys Cys Cys Leu
-20 -15 -10
Trp Lys Ile Ser Gly Cys Glu Glu Val Pro Leu Thr Tyr Asn Leu Leu
- 5
Lys Cys Leu Leu Asp Lys Ala His Cys Val Leu Leu Thr Pro Cys Gly
                 20
                                          25
       15
Tyr Ile Phe Ser Leu Ile Ser Pro Glu Ile Leu Lys Leu Thr Leu Ile
            35
 3 0
                                40
Thr Leu Xaa Ile Leu Leu Ile Leu Lys Asn Leu His Leu Leu Trp Leu
                           55
45 50
Thr Val Ser Ser Xaa Cys Val His Arg Ser Ser Ala Arg Lys Glu Lys
                             70
             65
<210> 476
<211> 41
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -24..-1
<400> 476
Met His Thr Phe Ala Asn Asp Arg Gly Leu Tyr Arg Ile Leu Leu
         -20 -15
His Phe Tyr Cys Leu Leu Arg Ser Ser Glu Tyr Ile Leu Gly Tyr Lys
       -5
                   1
```

<210> 477 <211> 113 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1

10

Val Leu Gly Val Phe Phe Pro Ile Leu

<400> 477 Met Arg Xaa Lys Trp Lys Met Gly Gly Met Lys Tyr Ile Phe Ser Leu -15 -20 Leu Phe Phe Leu Leu Glu Gly Gly Xaa Thr Glu Gln Val Xaa His - 5 Ser Glu Thr Tyr Cys Met Phe Gln Asp Lys Lys Tyr Arg Val Gly Glu 10 15 Arg Trp His Pro Tyr Leu Glu Pro Tyr Gly Leu Val Tyr Cys Val Asn . 30 Cys Ile Cys Ser Glu Asn Gly Asn Val Leu Cys Ser Arg Val Arg Cys 45 4.0 Pro Asn Val His Cys Leu Ser Pro Val His Ile Pro His Leu Cys Cys 60 Pro Arg Cys Pro Glu Asp Ser Leu Pro Pro Val Asn Asn Xaa Val Thr Ser

<210> 478 <211> 250 <212> PRT <213> Home sapieus

210

Asp Trp Ile Gln Glu Thr Met Lys Asn Asn

<220> <221> SIGNAL <222> -18..-1

<400> 478 Met Arg Ile Leu Gln Leu Ile Leu Leu Ala Leu Ala Thr Gly Leu Val -10 -5 -15 Gly Gly Glu Thr Arg Ile Ile Lys Gly Phe Glu Cys Lys Pro His Ser Gln Pro Trp Gln Ala Ala Leu Phe Glu Lys Thr Arg Leu Leu Cys Gly Ala Thr Leu Ile Ala Pro Arg Trp Leu Leu Thr Ala Ala His Cys Leu Lys Pro Arg Tyr Ile Xaa His Leu Gly Gln His Asn Leu Gln Lys Glu 55 Glu Gly Cys Glu Gln Thr Arg Thr Ala Thr Glu Ser Phe Pro His Pro 70 Gly Phe Asn Asn Ser Leu Pro Asn Lys Asp Xaa Xaa Asn Asp Ile Met 85 Leu Val Xaa Met Xaa Ser Pro Val Ser Ile Thr Trp Ala Val Arg Pro 105 100 Leu Thr Leu Ser Ser Arg Cys Val Thr Ala Gly Thr Ser Cys Leu Ile 115 120 Ser Gly Trp Gly Ser Thr Ser Ser Pro Gln Leu Arg Leu Pro His Thr 135 140 130 Leu Arg Cys Ala Asn Ile Thr Ile Ile Glu His Gln Lys Cys Glu Asn . 150 155 145 Ala Tyr Pro Gly Asn Ile Thr Asp Thr Met Val Cys Ala Ser Val Gln 165 170 Glu Gly Gly Lys Asp Ser Cys Gln Gly Asp Ser Gly Gly Pro Leu Val 180 185 Cys Asn Gln Ser Leu Gln Gly Ile Ile Ser Trp Gly Gln Asp Pro Cys 200 195 Ala Ile Thr Arg Lys Pro Gly Val Tyr Thr Lys Val Cys Lys Tyr Val 215 220

230

```
<210> 479
<211> 151
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 479
Met Ala Ala Ser Thr Ser Met Val Pro Val Ala Val Thr Ala Ala Val
                       -15
                                           -10
Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu Arg Glu Ile
Lys Lys Gln Leu Leu Leu Ile Ala Gly Leu Thr Arg Glu Arg Gly Leu
                              20
Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser Leu Pro Ala
                          35
Leu Pro Leu Ala Glu Leu Gln Pro Pro Pro Pro Ile Thr Glu Glu Asp
                       5.0
                                        5.5
Ala Gln Asp Met Asp Ala Tyr Thr Lev Ala Lys Ala Tyr Phe Asp Val
                                      7.0
                  65
Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys Asn Ala Arg
               80
                                  85
Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val Arg Ala Ile
                            100
           95
Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe Arg Thr Asn
       110
                          115
Gly Lys Val Lys Ser Phe Lys
<210> 480
<211> 239
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -25..-1
<400> 480
Met Pro Arg Lys Arg Lys Cys Asp Leu Arg Ala Val Arg Val Gly Leu
                   -20
                          -15
Leu Leu Gly Gly Gly Val Tyr Gly Ser Arg Phe Arg Phe Thr Phe
               -5
Pro Gly Cys Arg Ala Leu Ser Pro Trp Arg Val Arg Xaa Gln Arg Arg
                           15
Arg Cys Glu Met Ser Thr Met Phe Ala Asp Thr Leu Leu Ile Val Phe
Ile Ser Val Cys Thr Ala Leu Leu Ala Glu Gly Ile Thr Trp Val Leu
                   45
                                      50
Val Tyr Arg Thr Asp Lys Tyr Lys Arg Leu Lys Ala Glu Val Glu Lys
                                  65
Gln Ser Lys Lys Leu Glu Lys Lys Glu Thr Ile Thr Glu Ser Ala
           75
                              80
Gly Arg Gin Gln Lys Lys Ile Glu Arg Xaa Xaa Xaa Xaa Leu Xaa
                           95
```

Asn Asn Asn Arg Asp Leu Ser Met Val Arg Met Lys Ser Met Phe Ala 110 Ile Gly Phe Cys Phe Thr Ala Leu Met Gly Met Phe Asn Ser Ile Phe 125 130 Asp Gly Arg Val Val Ala Lys Leu Pro Phe Thr Pro Leu Ser Xaa Xaa 140 145 Xaa Gly Leu Ser His Arg Asn Leu Leu Gly Asp Asp Thr Thr Asp Cys 160 Ser Phe Ile Phe Leu Xaa Ile Leu Cys Thr Met Ser Ile Arg Gln Asn 175 Ile Gln Lys Ile Leu Gly Leu Ala Pro Ser Arg Ala Ala Thr Lys Gln 190 195 Ala Gly Gly Phe Leu Gly Pro Pro Pro Pro Ser Gly Lys Phe Ser 205

<210> 481 <211> 208 <212> PRT <213> Homo sapiens <220> ~221> SIGNAL <222> -92..-1 <400> 481 Met Arg Glu Pro Gln Lys Arg Thr Ala Thr Ile Ala Lys Xaa Xaa Ala -85 Xaa Glu Gly Leu Arg Asp Pro Tyr Gly Arg Leu Cys Gly Ser Glu His -70 -65 Pro Arg Arg Pro Pro Glu Arg Pro Glu Glu Asp Pro Ser Thr Pro Glu -55 -50 Glu Ala Ser Thr Thr Pro Glu Glu Ala Ser Ser Thr Ala Gln Ala Gln -35 Lys Pro Ser Val Pro Arg Ser Asn Phe Gln Gly Thr Lys Lys Ser Leu -20 -15 Leu Met Ser Ile Leu Ala Leu Ile Phe Ile Met Gly Asn Ser Ala Lys - 5 Glu Ala Leu Val Trp Lys Val Leu Gly Lys Leu Gly Met Gln Pro Gly 10 15 Arg Xaa His Ser Ile Phe Gly Asp Pro Lys Lys Ile Val Thr Glu Xaa 30 25 Phe Val Arg Arg Gly Tyr Leu Ile Tyr Xaa Pro Val Pro Arg Xaa Ser 40 45 Pro Val Glu Tyr Xaa Phe Phe Trp Gly Pro Arg Ala His Val Glu Ser 60 Ser Xaa Leu Lys Xaa Xaa His Phe Val Ala Arg Val Arg Asn Arg Cys 75 Ser Lys Asp Trp Pro Cys Asn Tyr Asp Trp Asp Ser Asp Asp Ala 90 Glu Val Glu Ala Ile Leu Asn Ser Gly Ala Xaa Gly Tyr Ser Ala Pro

<210> 482 <211> 86 <212> PRT <213> Homo sapiens <221> SIGNAL <222> -39..-1

<400> 482

 Met Asn Val
 Gly Thr Ala His Xaa Xaa Val Asn Pro Asn Thr Arg Val -35

 Met Asn Ser Arg Gly Ile Trp Leu Ser Tyr Val Leu Ala Ile Gly Leu -10

 Leu His Ile Val Leu Leu Ser Ile Pro Phe Val Ser Val Pro Val Val -5

 Trp Thr Leu Thr Asn Leu Ile His Asn Met Gly Met Tyr Ile Phe Leu 20

 His Thr Val Lys Gly Thr Pro Phe Glu Thr Pro Asp Gln Gly Lys Ala 30

 Arg Leu Leu Thr His Trp

Arg Leu Leu Thr His Trp
45

<210> 483

<211> 40

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -27..-1

<400> 483

Met Arg Thr Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr
-25

Leu Leu Leu Leu Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly
-10

Leu Ser Leu Arg Ser Ala Met Ser

10

<210> 484

<211> 65

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -16..-1

<400> 484

 Met
 Leu
 Gly
 Phe
 Leu
 Phe
 Leu
 Ser
 Phe
 Val
 Leu
 Met
 Tyr
 Asp
 Gly

 Leu
 Arg
 Leu
 Phe
 Gly
 Ile
 Leu
 Ser
 Thr
 Cys
 Arg
 Val
 His
 His
 Thr
 Met

 1
 5
 10
 15

 Asn
 Gln
 Phe
 Leu
 Ile
 Asp
 Ile
 Ser
 Ser
 Phe
 Thr
 Ser
 Arg
 Val
 Lys
 Lys

 Lys
 1le
 Phe
 Leu
 Phe
 Tyr
 Ala
 Phe
 Xaa
 Gly
 Cys
 Xaa
 Phe
 Gln
 Ser
 Ala

Thr

<210> 485

<211> 130

PCT/IB98/02122 -

```
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -55..-1
<400> 485
Met Ala Met Trp Asn Arg Pro Xaa Xaa Xaa Leu Pro Gln Gln Pro Leu
                         -45
Xaa Ala Glu Pro Thr Ala Glu Gly Glu Pro His Leu Pro Thr Gly Arg
                                                 -25
                                 -30
Xaa Xaa Thr Glu Ala Asn Arg Phe Ala Tyr Ala Ala Leu Cys Gly Ile
                           -15 -10
Ser Leu Ser Gln Leu Phe Pro Glu Pro Glu His Ser Ser Phe Cys Thr
                         1
Glu Phe Met Ala Gly Leu Val Xaa Trp Leu Glu Leu Ser Glu Ala Val
           15
                                 20
Leu Pro Thr Met Thr Ala Phe Ala Ser Gly Leu Gly Gly Glu Gly Xaa
            30
                                 35
Xaa Cys Val Cys Ser Asn Phe Thr Glu Gly Pro His Leu Glu Gly Arg
                      50
       4.5
Pro Asp Gly Asp His Ser Gly Pro Ser Glu Leu Leu Thr Gln Gly Trp
              65
Ala Leu
  75
<210> 486
<211> 209
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -84..-1
<400> 486
Met Val Asn Phe Pro Gln Lys Ile Ala Gly Glu Leu Tyr Gly Pro Leu
              -80
                                  -75
Met Leu Val Phe Thr Leu Val Ala Ile Leu Leu His Gly Met Lys Thr
                              -60
                                                 -55
Ser Asp Thr Ile Ile Arg Glu Gly Thr Leu Met Gly Thr Ala Ile Gly
                          -45
                                             -40
Thr Cys Phe Gly Tyr Trp Leu Gly Val Ser Ser Phe Ile Tyr Phe Leu
                     -30
                                        -25
Ala Tyr Leu Cys Asn Ala Gln Ile Thr Met Leu Gln Met Leu Ala Leu
                                     -10
                  -15
Leu Gly Tyr Gly Leu Phe Gly His Cys Ile Val Leu Phe Ile Thr Tyr
Asn Ile His Leu Arg Ala Leu Phe Tyr Leu Phe Trp Leu Leu Val Gly
                          20
Gly Leu Ser Thr Leu Arg Met Val Ala Val Leu Val Ser Arg Thr Val
                      35
                                         40
Gly Pro Thr Xaa Arg Xaa Leu Leu Cys Gly Thr Leu Ala Ala Leu His
                  50
                                      55
Met Leu Phe Leu Leu Tyr Leu His Phe Ala Tyr His Lys Xaa Val Xaa
                                  70
              65
Gly Ile Leu Asp Thr Leu Glu Gly Pro Asn Ile Pro Pro Ile Gln Arg
                              85
```

Val Pro Arg Asp Ile Pro Ala Met Leu Pro Ala Ala Arg Leu Pro Thr

```
100
       95
                                             105
Thr Val Leu Asn Ala Thr Ala Lys Ala Val Ala Val Thr Leu Gln Ser
            115
                                 120
His
125
<210> 487
<211> 36
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -17..-1
<400> 487
Met Gly Trp Gln Arg Trp Trp Cys Phe His Leu Gln Ala Glu Ala Ser
    -15 -10
Ala His Pro Pro Gln Gly Leu Gln Ala Gln Phe Ser Cys Cys Pro Trp
                                     10
Val Gly Ile Cys
<210> 488
<211> 44
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -29..-1
<400> 488
Met Met Ser Ser Glu Leu Arg Arg Asn Pro His Phe Leu Lys Ser Asn
                         -20
              -25
Leu Phe Leu Gln Leu Leu Val Ser His Glu Ile Val Cys Ala Thr Glu
Thr Val Thr Thr Asn Phe Leu Arg His Glu Lys Ala
 5
                      10
<210> 489
<211> 163
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -52..-1
<400> 489
Met Glu His Tyr Arg Lys Ala Gly Ser Val Glu Leu Pro Ala Pro Ser
    -50
                          -45
                                             -40
Pro Met Pro Gln Leu Pro Pro Asp Thr Leu Glu Met Arg Val Arg Asp
  -35
                      -30
                                         -25
Gly Ser Lys Ile Arg Asn Leu Leu Gly Leu Ala Leu Gly Arg Leu Glu
                  -15
                                    -10
Gly Gly Ser Ala Arg His Val Val Phe Ser Gly Ser Gly Arg Ala Ala
```

```
Gly Lys Ala Val Ser Cys Ala Glu Ile Val Lys Arg Arg Val Pro Gly
                           20
Leu His Gln Leu Thr Lys Leu Xaa Phe Leu Gln Thr Glu Asp Ser Trp
                       35
Val Pro Xaa Ser Pro Asp Thr Gly Leu Xaa Pro Leu Thr Val Arg Arg
                                       55
His Val Pro Ala Xaa Trp Val Leu Leu Xaa Arg Asp Pro Leu Asp Pro
                                   70
Asn Glu Cys Gly Tyr Gln Pro Pro Gly Ala Pro Pro Gly Leu Gly Ser
                               85
Met Pro Ser Ser Ser Cys Gly Pro Arg Ser Xaa Lys Arg Ala Xaa Xaa
                          100
Thr Arg Ser
  110
<210> 490
<211> 64
<212> PRT
<213> Homo sapiens
₹220 \
<221> SIGNAL
<222> -47..-1
<400> 490
Met His Gly Phe Glu Ile Ile Ser Leu Lys Glu Glu Ser Pro Leu Gly
                    -40
                                      -35
Lys Val Ser Gln Gly Pro Leu Phe Asn Val Thr Ser Gly Ser Ser Ser
                       -25
Pro Val Thr Trp Leu Gly Leu Leu Ser Phe Gln Asn Leu His Cys Phe
-15
                   -10
Pro Asp Leu Pro Thr Glu Met Pro Leu Xaa Ala Lys Gly Xaa Asn Thr
                            10
<210> 491
<211> 218
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -50..-1
<400> 491
Met His His Gly Leu Thr Pro Leu Leu Gly Val His Glu Gln Lys
                                      -40
Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys Ala Asn Leu Asn Ala
               ~30
                                  -25
Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu Ala Val Cys Cys Gly
           -15
                              -10
Ser Ala Ser Ile Val Ser Leu Leu Glu Gln Asn Ile Asp Val Ser
Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys Tyr Ala Val Ser Ser
                  20
                                     25
Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp Tyr Lys Xaa Lys Gln
               35
                                  40
Xaa Leu Lys Val Ser Ser Glu Asn Ser Asn Pro Xaa Gln Asp Leu Lys
```

```
55
          50
Leu Thr Ser Glu Glu Glu Ser Gln Arg Leu Lys Gly Ser Glu Asn Ser
                          70
Gln Pro Glu Glu Met Ser Gln Glu Pro Glu Ile Asn Xaa Gly Gly Asp
                      85
Arg Lys Val Glu Xaa Xaa Met Lys Lys His Gly Ser Xaa His Met Gly
                                    105
                 100
Phe Pro Xaa Asn Leu Xaa Asn Gly Ala Thr Ala Asp Asn Gly Asp Asp
                                120
             115
Gly Leu Ile Pro Pro Xaa Lys Xaa Xaa Thr Pro Glu Ser Xaa Gln Phe
                            135
          130
Pro Asp Thr Glu Asn Glu Gln Tyr His Arg Asp Phe Ser Gly His Pro
      145 150
Xaa Phe Pro Thr Thr Leu Pro Ile Lys Gln
                    165
  160
```

<210> 492 <211> 216 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1

<400> 492 Met Val Cys Val Leu Val Leu Ala Ala Ala Ala Gly Ala Val Ala Val -10 Phe Leu Ile Leu Arg Ile Trp Val Val Leu Arg Ser Met Asp Val Thr 10 Pro Arg Glu Ser Leu Ser Ile Leu Val Val Ala Gly Ser Gly Gly His 25 20 Thr Thr Glu Ile Leu Arg Leu Leu Gly Ser Leu Ser Asn Ala Tyr Ser 40 Pro Arg His Tyr Val Ile Ala Asp Thr Asp Glu Met Ser Ala Asn Lys 60 55 Ile Asn Ser Phe Glu Leu Xaa Arg Xaa Asp Arg Xaa Pro Ser Asn Met 75 70 Xaa Thr Lys Tyr Tyr Ile His Arg Ile Pro Xaa Ser Arg Glu Val Gln 90 85 Gln Ser Trp Pro Ser Thr Val Xaa Thr Thr Leu His Ser Met Trp Leu 100 105 Ser Xaa Pro Leu Ile His Arg Val Lys Pro Xaa Leu Val Leu Cys Asn 125 120 Gly Pro Gly Thr Cys Val Pro Ile Cys Val Ser Ala Leu Leu Leu Gly 135 140 Ile Leu Gly Ile Lys Lys Val Ile Ile Val Tyr Val Glu Ser Ile Cys 155 150 Arg Val Lys Thr Leu Ser Met Ser Gly Lys Ile Leu Phe His Leu Ser 170 175 Asn Tyr Phe Ile Val Gln Trp Pro Ala Leu Lys Glu Lys Tyr Pro Lys 185 Ser Val Tyr Leu Gly Arg Ile Val

200

<210> 493 <211> 134 <212> PRT <213> Homo sapiens

<220>

<221> SIGNAL

<222> -19..-1

<400> 493

Met Pro Leu Gly Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala Gly
-15 -10 -5

-347-

Gly Phe Ala Ile Val Tyr Tyr Leu Ile Gln Lys Phe His Ser Arg Thr

Leu Tyr Tyr Lys Leu Ala Val Glu Gln Leu Gln Xaa His Pro Glu Ala
15 20 25

Gln Glu Ala Leu Gly Pro Pro Leu Asn Ile His Tyr Leu Lys Leu Ile
30 35 40 45

30 35 40 45
Asp Arg Glu Asn Phe Val Asp Ile Val Xaa Ala Lys Leu Lys Ile Pro

Val Ser Gly Ser Lys Ser Glu Gly Leu Leu Tyr Val His Ser Ser Arg

65 70 75

Gly Gly Pro Phe Gln Arg Trp His Leu Asp Glu Val Phe Leu Glu Leu 80 85 90

Lys Asp Gly Gln Gln Ile Pro Val Phe Lys Leu Ser Gly Glu Asn Gly 95 100 105

Asp Glu Val bys bys Clu 110 115

110

<210> 494

<211> 85

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -16..-1

<400> 494

Met Ala Val Thr Ala Leu Ala Ala Xaa Thr Trp Leu Gly Val Trp Gly
-15 -5

Val Arg Thr Met Gln Ala Arg Gly Phe Gly Ser Asp Gln Ser Glu Asn
1 5 10 15

Val Asp Arg Gly Ala Gly Ser Ile Arg Glu Ala Gly Gly Ala Phe Gly
20 25 30

Lys Arg Glu Gln Ala Glu Glu Glu Arg Tyr Phe Arg Ala Gln Ser Thr

Glu Gln Leu Ala Xaa Leu Lys Lys Xaa His Glu Glu Glu Ile Val His 50 55 60

His Arg Glu Gly Asp

65

<210> 495

<211> 292

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -29..-1

```
<400> 495
Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn His Thr Phe
                                   -20
Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr Glu Tyr Thr
           -10
                               - 5
Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser Leu His Tyr
                       10
Leu Leu Pro Tyr Leu Leu Gly Val Asn Leu Phe Phe Thr
                   25
                                       30
Leu Thr Cys Gly Thr Asn Pro Gly Ile Ile Thr Lys Ala Asn Glu Leu
               40
Leu Phe Leu His Val Tyr Glu Phe Asp Glu Xaa Met Phe Pro Lys Asn
                              60
Val Arg Cys Ser Thr Cys Asp Leu Arg Lys Pro Ala Arg Ser Xaa His
                           75
Cys Xaa Val Cys Asn Trp Cys Val His Arg Phe Xaa His His Cys Val
                       90
Trp Val Asn Asn Cys Ile Gly Ala Trp Asn Ile Arg Xaa Phe Leu Ile
                   105
                                      110
Tyr Val Leu Thr Leu Thr Ala Ser Ala Ala Thr Val Ala Ile Val Ser
               120
                                   125
Thr Thr Phe Leu Val His Leu Val Val Met Ser Asp Leu Tyr Gln Glu
                               140
                                         145
Thr Tyr Ile Asp Asp Lau Bly His Leu His Val Met Asp Thr Val Phe
       150
                           155
                                              160
Leu Ile Gln Tyr Leu Phe Leu Thr Phe Pro Arg Ile Val Phe Met Leu
                       170
                                          175
Gly Phe Val Val Leu Xaa Phe Leu Leu Gly Gly Tyr Leu Leu Phe
                   185
                                       190
Val Leu Tyr Leu Ala Ala Thr Asn Gln Thr Thr Asn Glu Trp Tyr Arg
               200
                                   205
Xaa Asp Trp Ala Trp Cys Gln Arg Cys Pro Leu Val Ala Trp Pro Pro
                               220
                                                  225
Ser Ala Glu Pro Gln Val His Arg Asn Ile His Ser His Gly Leu Arg
                          235
Xaa Asn Leu Gln Glu Ile Phe Leu Pro Ala Phe Pro Cys His Glu Arg
                      250
Lys Lys Gln Glu
260
```

<210> 496 <211> 122 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -56..-1

<400> 496

 Met
 Thr
 Gly
 Phe
 Leu
 Leu
 Pro
 Pro
 Ala
 Ser
 Arg
 Gly
 Thr
 Arg
 Arg
 Arg
 Ser
 -45

 Cys
 Ser
 Arg
 Ser
 Arg
 Gln
 Thr
 Arg
 Arg

<210> 497 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 497 Met Leu Xaa Leu Ser Arg Ala Thr Lys Xaa Gly Arg Ala Arg Trp Leu -20 Met Pro Val Ile Pro Ala Leu Gln Glu Ala Xaa Ala Gly Gly Ser Arg -10 - 5 Gly Gln Glu Phe Glu Thr Str Teu Ala Ash Met Glu Thr Glu Ala Gly 10 15 Glu Leu Leu Lys Pro Arg Arg Arg Leu Gln

<210> 498 <211> 99 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -13..-1

<210> 499 <211> 99 <212> PRT <213> Homo sapiens

```
<220>
<221> SIGNAL
<222> -13..-1
<400> 499
Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg Arg Pro
                                - 5
           -10
Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu Ala His
                                           15
                       10
Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys Trp Arg
                                       30
                   25
Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn Ser Ser
                                   45
               40
Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr Pro Met
                               60
Arg Arg Ser Ser Cys His Leu Xaa Cys Gln Val Ile Phe Leu Leu Gly
Arg Gln Leu
  85
<210> 500
<211> 108
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -25..-1
<400> 500
Met Ser Leu Thr Ser Ser Ser Ser Val Arg Val Glu Trp Ile Ala Ala
           -20
                                       -15
Val Thr Ile Ala Ala Gly Thr Ala Ala Ile Gly Tyr Leu Ala Tyr Lys
               - 5
Arg Phe Tyr Val Lys Asp His Arg Asn Lys Ala Met Ile Asn Leu His
     10
Ile Gln Lys Asp Asn Pro Lys Ile Val His Ala Phe Asp Met Glu Asp
                       30
Leu Gly Asp Lys Ala Val Tyr Cys Arg Cys Trp Arg Ser Lys Lys Phe
                                       50
Pro Phe Cys Asp Gly Ala His Thr Lys His Asn Glu Glu Thr Gly Asp
Asn Val Gly Pro Leu Ile Ile Lys Lys Glu Thr
                               8.0
<210> 501
<211> 183
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -15..-1
<400> 501
```

Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu Ala Val Leu Ala Trp
-15 -10 -5 1
Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg Met Lys Ser Arg Glu

WO 99/31236

```
10
Gln Gly Arg Arg Leu Gly Ala Glu Ser Arg Thr Leu Leu Val Ile Ala
                           25
His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro Thr Val Leu Gly Leu
                       40
Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys Phe Ser Ala Gly Asn
                                     60
                  55
Tyr Tyr Asn Gln Gly Glu Thr Arg Lys Lys Glu Leu Leu Gln Ser Cys
                                  75
Asp Val Leu Gly Ile Pro Leu Ser Ser Val Met Ile Ile Asp Asn Arg
                              90
Asp Phe Pro Xaa Asp Pro Gly Met Gln Trp Asp Thr Xaa His Val Ala
                          105
                                          110
Xaa Val Leu Leu Gln His Ile Glu Val Asn Gly Ile Asn Leu Val Val
                      120
                                         125
Thr Phe Asp Ala Gly Gly Xaa Ser Gly His Ser Asn His Ile Ala Leu
                  135
                               140
Tyr Ala Ala Val Arg Lys Leu Glu Gly Gln Ile Cys Lys Pro Cys Gly
                                 155
              150
Thr Gly Gln Asp Phe Lys Glu
```

<210 > 502 <211 > 98 <212 > PRT <213 > Homo sapiens <220 > <221 > SIGNAL <222 > -15..-1 <400 > 502 Met Glu Ala Met Trp Letters -15 Gly Phe Leu Trp Val Try

 Met
 Glu
 Ala
 Met
 Trp
 Leu
 Leu
 Cys
 Val
 Ala
 Leu
 Ala
 Val
 Val
 Leu
 Ala
 Trp
 Val
 Trp
 Asp
 Ser
 Ser
 Glu
 Arg
 Met
 Lys
 Ser
 Arg
 Glu

 Glu
 Phe
 Leu
 Trp
 Asp
 Asp
 Leu
 Glu
 Asp
 Arg
 Arg

<210> 503 <211> 183 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -57..-1

<400> 503
Met Asp Val Thr Gly Asp Glu Glu Glu Glu Ile Lys Gln Glu Ile Asn
-55
-50
-45

```
Met Leu Lys Lys Tyr Ser His His Arg Asn Ile Ala Thr Tyr Tyr Gly
                        -35
Ala Phe Ile Lys Lys Asn Pro Pro Gly Met Asp Asp Gln Leu Trp Leu
                                        -15
Val Met Glu Phe Cys Gly Ala Gly Ser Val Thr Asp Leu Ile Lys Asn
Thr Lys Gly Asn Thr Leu Lys Glu Glu Trp Ile Ala Tyr Ile Cys Xaa
                           15
Glu Ile Leu Arg Gly Leu Xaa His Leu His Gln His Lys Val Ile His
                        30
Arg Xaa Ile Lys Gly Gln Asn Val Leu Leu Thr Glu Asn Ala Glu Val
                    45
                                       50
Lys Leu Val Asp Phe Gly Xaa Xaa Ala Gln Leu Asp Arg Thr Val Gly
                60
                                    65
Arg Xaa Asn Thr Phe Ile Gly Thr Pro Tyr Trp Met Ala Pro Xaa Val
                               80
Ile Ala Cys Asp Glu Asn Pro Xaa Ala Thr Tyr Asp Phe Lys Xaa Asp
                           95
Leu Trp Ser Leu Gly Ile Thr Ala Ile Glu Met Ala Glu Gly Leu Pro
Leu Ser Val Thr Cys Thr Pro
```

```
<210> 504
<211> 140
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -14..-1
<400> 504
Met Phe Leu Thr Ala Leu Leu Trp Arg Gly Arg Ile Pro Gly Arg Gln
                -10
Trp Ile Gly Lys His Arg Arg Pro Arg Phe Val Ser Leu Arg Ala Lys
Gln Asn Met Ile Arg Arg Leu Glu Ile Glu Ala Glu Asn His Tyr Trp
                        25
Leu Ser Met Pro Tyr Met Thr Arg Glu Gln Glu Arg Gly His Ala Ala
                    40
Leu Arg Arg Arg Glu Ala Phe Glu Ala Ile Lys Ala Ala Ala Thr Ser
                55
                                    60
Lys Phe Pro Pro His Arg Phe Ile Ala Asp Gln Leu Asp His Leu Asn
                                75
Xaa His Gln Glu Met Val Leu Ile Leu Ser Arg His Pro Trp Ile Leu
                           .90
Trp Ile Thr Glu Leu Thr Ile Phe Thr Trp Ser Gly Leu Lys Asn Cys
                       105
                                            110
Ser Leu Cys Glu Asn Glu Leu Trp Thr Ser Leu Tyr
```

120

<210> 505 <211> 59 <212> PRT <213> Homo sapiens

<221> SIGNAL <222> -14..-1

<400> 505

Met Ala Ala Leu Val Thr Val Leu Phe Thr Gly Val Arg Arg Leu His
-10 -5 1

. ....

Cys Ser Ala Xaa Leu Gly Arg Ala Ala Ser Gly Xaa Tyr Ser Arg Asn 5 10 15

Trp Leu Pro Thr Pro Pro Ala Thr Gly Pro Leu Pro Ser Ser Gln Thr
20 25 30

Gly His Met Arg Met Ala Ala Leu Leu Pro Gln 35 40 45

<210> 506

<211> 101

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -36..-1

4400> 506

Met Gly Pro Tyr Asn Val Ala Val Pro Ser Asp Val Ser His Ala Arg
-35
-30
-25

Phe Tyr Phe Leu Phe His Arg Pro Leu Arg Leu Leu Asn Leu Leu Ile
-20 -15 -10 -5

Leu Ile Glu Gly Ser Val Val Phe Tyr Gln Leu Tyr Ser Leu Leu Arg 1 5 5 10

Ser Glu Lys Trp Asn His Thr Leu Ser Met Ala Leu Ile Leu Phe Cys
15 20 25

Asn Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Arg Xaa Xaa Leu Gly 30 40

Arg Ala Tyr Ser Tyr Pro Leu Asn Ser Tyr Glu Leu Lys Ala Asn Xaa 45 50 55 60

Ala Ala Ser Xaa Gln

65

<210> 507

<211> 341

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -55..-1

<400> 507

Met Arg Lys Val Val Leu Ile Thr Gly Ala Ser Ser Gly Ile Gly Leu
-55 -50 -45 -45

Ala Leu Cys Lys Arg Leu Leu Ala Glu Asp Asp Glu Leu His Leu Cys
-35 -30 -25

Leu Ala Cys Arg Asn Met Ser Lys Ala Glu Ala Val Cys Ala Ala Leu
-20 -15 -10

Leu Ala Ser His Pro Thr Ala Glu Val Thr Ile Val Gln Val Asp Val -5 1 5

Ser Asm Leu Gln Ser Phe Phe Arg Ala Ser Lys Glu Leu Lys Gln Arg
10 15 20 25

Phe Gln Arg Leu Asp Cys Ile Tyr Leu Asn Ala Gly Ile Met Pro Asn 35 Pro Gln Leu Asn Ile Lys Ala Leu Phe Phe Gly Leu Phe Ser Arg Lys 50 Val Ile His Met Phe Ser Thr Ala Glu Gly Leu Leu Thr Gln Gly Asp 65 Lys Ile Thr Ala Asp Gly Leu Gln Glu Val Phe Glu Thr Asn Val Phe 80 Gly His Phe Ile Leu Ile Arg Glu Leu Glu Pro Leu Leu Cys His Ser 100 95 Asp Asn Pro Ser Gln Leu Ile Trp Thr Ser Ser Arg Ser Ala Arg Lys 115 110 Ser Asn Phe Ser Leu Glu Asp Phe Gln His Ser Lys Gly Lys Glu Pro 130 Tyr Ser Ser Ser Lys Tyr Ala Thr Asp Leu Leu Ser Val Ala Leu Asn 145 Arg Asn Phe Asn Gln Gln Gly Leu Tyr Ser Asn Val Ala Cys Pro Gly 165 160 Thr Ala Leu Thr Asn Leu Thr Tyr Gly Ile Leu Pro Pro Phe Ile Trp 180 175 Thr Leu Leu Met Pro Ala Ile Leu Leu Leu Arg Phe Phe Ala Asn Ala 190 195 Phe Thr Leu Thr Pro Tyr Asn Gly Thr Glu Ala Leu Val Trp Leu Phe 210 His Gln Lys Pro Glu Ser Leu Asn Pro Leu Ile Lys Tyr Leu Ser Ala 220 225 Thr Thr Gly Phe Gly Arg Asn Tyr Ile Met Thr Gln Lys Met Asp Leu 245 240 Asp Glu Asp Thr Ala Glu Lys Phe Tyr Gln Lys Leu Leu Glu Leu Glu 255 260 Lys His Ile Arg Val Thr Ile Gln Lys Thr Asp Asn Gln Ala Arg Leu 270 Ser Gly Ser Cys Leu 285

<210> 508 <211> 108 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -42..-1

```
<210> 509
<211> 80
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -26..-1
<400> 509
Met Glu Glu Ile Ser Ser Pro Leu Val Glu Phe Val Lys Val Leu Cys
                    -20
Thr Asn Gln Val Leu Ile Thr Ala Arg Ala Val Pro Thr Lys Lys Ala
Ser Val Arg Cys Val Glu Lys Arg Phe Trp Ile Pro Lys Thr Thr Ser
Lys His Leu Ser Arg Cys Ile Asp Gly Ile Ser Gly Phe Leu Asn Asp
                           30
Phe Thr Phe Cys Leu Glu Phe Ser Arg His Arg Cys Gln Leu Thr Glu
            45
<210> 510
<211> 158
```

<212> PRT <213> Homo sapiens <221> SIGNAL <222> -44..-1 <400> 510 Met Ala Gly Phe Leu Asp Asn Phe Arg Trp Pro Glu Cys Glu Cys Ile -40 -35 Asp Trp Ser Glu Arg Arg Asn Ala Val Ala Ser Val Val Ala Gly Ile -25 -20 Leu Phe Phe Thr Gly Trp Trp Ile Met Ile Asp Ala Ala Val Val Tyr -5 Pro Lys Pro Glu Gln Leu Asn His Ala Phe His Thr Cys Gly Val Phe 10 15 Ser Thr Leu Ala Phe Phe Met Ile Asn Ala Val Ser Asn Ala Gln Val 30 25 Arg Gly Asp Ser Tyr Glu Ser Gly Cys Leu Gly Arg Thr Gly Ala Arg 45 Val Trp Leu Phe Ile Gly Phe Met Leu Met Phe Gly Ser Leu Ile Ala 60 Ser Met Trp Ile Leu Phe Gly Ala Tyr Val Thr Gln Asn Thr Asp Val

Tyr Pro Gly Leu Ala Val Phe Phe Gln Asn Ala Leu Ile Phe Phe Ser

Thr Leu Ile Tyr Lys Phe Gly Arg Thr Glu Glu Leu Trp Thr

90

105

<210> 511 <211> 130 <212> PRT <213> Homo sapiens

PCT/IB98/02122

<220> <221> SIGNAL <222> -28..-1 <400> 511 Met Asn Trp Glu Leu Leu Trp Leu Leu Val Leu Cys Ala Leu Leu -20 -15 -25 Leu Leu Leu Val Gln Leu Leu Arg Phe Leu Arg Ala Asp Gly Asp Leu Thr Leu Leu Trp Ala Glu Trp Gln Gly Arg Arg Pro Glu Trp Glu Leu 10 Thr Asp Met Val Val Trp Val Thr Gly Ala Ser Ser Gly Ile Gly Glu Glu Leu Ala Tyr Gln Leu Ser Lys Leu Gly Val Ser Leu Val Leu Ser Ala Arg Arg Val His Glu Leu Glu Arg Val Lys Arg Arg Cys Leu Glu 60 Asn Gly Asn Leu Lys Glu Lys Asp Ile Leu Val Leu Pro Leu Asp Leu Thr Asp Thr Gly Ser His Glu Ser Gly Tyr Gln Ser Cys Ser Pro Gly 90 Ile Trp <210> 512 <211> 199 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -62..-1 <400> 512 Met Ser Gln Arg Ser Leu Cys Met Asp Thr Ser Leu Asp Val Tyr Arg -55 -50 Xaa Leu Ile Glu Leu Asn Tyr Leu Gly Thr Val Ser Leu Thr Lys Cys -40 -35 Val Leu Pro His Met Ile Glu Arg Lys Gln Gly Lys Ile Val Thr Val -25 -20 Asn Ser Ile Leu Gly Ile Ile Ser Val Pro Leu Ser Ile Gly Tyr Cys -10 -5 Ala Ser Lys His Ala Leu Arg Gly Phe Phe Asn Gly Leu Arg Thr Glu 10 Leu Ala Thr Tyr Pro Gly Ile Ile Val Ser Asn Ile Cys Pro Gly Pro 25 Val Gln Ser Asn Ile Val Glu Asn Ser Leu Ala Gly Glu Val Thr Lys 40 45 Thr Ile Gly Asn Asn Gly Asn Gln Ser His Lys Met Thr Thr Ser Arg 60 Cys Val Arg Leu Met Leu Ile Ser Met Ala Asn Asp Leu Lys Glu Val 70 75 Trp Ile Ser Glu Gln Pro Phe Leu Leu Val Thr Tyr Leu Trp Gln Tyr

90

105

120

Ile Phe Lys Thr Lys His Asp

135

Met Pro Thr Trp Ala Trp Trp Ile Thr Asn Lys Met Gly Lys Lys Arg

Ile Glu Asn Phe Lys Ser Gly Val Asp Ala Xaa Ser Ser Tyr Phe Lys

110

125

```
<210> 513
<211> 180
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -25..-1
<400> 513
Met Asn Thr Val Leu Ser Arg Ala Asn Ser Leu Phe Ala Phe Ser Leu
           -20
                                    -15
Ser Val Met Ala Ala Leu Thr Phe Gly Cys Phe Ile Xaa Thr Ala Phe
       · -5
Lys Asp Arg Ser Val Pro Val Arg Leu His Val Ser Arg Ile Met Leu
   10
                         15
Lys Asn Val Glu Asp Phe Thr Gly Pro Arg Glu Arg Ser Asp Leu Gly
            30
Phe Ile Thr Phe Asp Ile Thr Ala Asp Leu Glu Asn Ile Phe Asp Trp
                 45
                                  50
Asn Val Lys Gin Leu Phe Leu Tyr Leu Ser Ala Glu Tyr Ser Thr Lys
            60
Asn Asn Ala Leu Asn Gln Xaa Val Leu Trp Asp Lys Ile Val Leu Arg
                            80
Gly Asp Asn Pro Lys Leu Leu Lys Asp Met Lys Thr Lys Tyr Phe
                               .. 100
                         95
Phe Phe Asp Asp Gly Asn Gly Leu Xaa Gly Asn Arg Asn Val Thr Leu
                     110
                                        115
Thr Leu Ser Trp Asn Val Val Pro Asn Ala Gly Ile Leu Pro Leu Val
                 125
                       130 135
Thr Gly Ser Gly His Val Ser Val Pro Phe Pro Asp Thr Tyr Glu Ile
                             145
Thr Lys Ser Tyr
          155
<210> 514
<211> 120
<212> PRT
<213> Bos taurus
<400> 514
Met Met Thr Gly Arg Gln Gly Arg Ala Thr Phe Gln Phe Leu Pro Asp
Glu Ala Arg Ser Leu Pro Pro Pro Lys Leu Thr Asp Pro Arg Leu Ala
                            25
Phe Val Gly Phe Leu Gly Tyr Cys Ser Gly Leu Ile Asp Asn Ala Ile
                         40
Arg Arg Arg Pro Val Leu Leu Ala Gly Leu His Arg Gln Leu Leu Tyr
                     55
Ile Thr Ser Phe Val Phe Val Gly Tyr Tyr Leu Leu Lys Arg Gln Asp
                  70
                                    75
Tyr Met Tyr Ala Val Arg Asp His Asp Met Phe Ser Tyr Ile Lys Ser
              85
```

90

His Pro Glu Asp Phe Pro Glu Lys Asp Lys Lys Thr Tyr Gly Glu Val

105

100

Phe Glu Glu Phe His Pro Val Arg

```
<210> 515
<211> 1082
<212> DNA
<213> Homo sapiens
<400> 515
gatcccagac ctcggcttgc agtagtgtta gactgaagat aaagtaagtg ctgtttgggc
                                                                  60
taacaggatc teetettgea gtetgeagee caggacgetg attecageag egeettaceg
                                                                 120
cgcagcccga agattcacta tggtgaaaat cgccttcaat acccctaccg ccgtgcaaaa
                                                                 180
ggaggaggcg cggcaagacg tggaggccct cctgagccgc acggtcagaa ctcagatact
                                                                 240
                                                                 300
gaccggcaag gagctccgag ttgccaccca ggaaaaagag ggctcctctg ggagatgtat
gcttactctc ttaggccttt cattcatctt ggcaggactt attgttggtg gagcctgcat
                                                                 360
                                                                 420
ttacaagtac ttcatgccca agagcaccat ttaccgtgga gagatgtgct tttttgattc
tgaggatect geaaatteee ttegtggagg agageetaac tteetgeetg tgaetgagga
                                                                 480
ggctgacatt cgtgaggatg acaacattgc aatcattgat gtgcctgtcc ccagtttctc
                                                                 540
tgatagtgac cctgcagcaa ttattcatga ctttgaaaag ggaatgactg cttacctgga
                                                                 600
cttgttgctg gggaactgct atctgatgcc cctcaatact tctattgtta tgcctccaaa
                                                                 660
aaatctggta gagctctttg gcaaactggc gagtggcaga tatctgcctc aaacttatgt
                                                                 720
                                                                 780
ggttcgagaa gacctagttg ctgtggagga aattcgtgat gttagtaacc ttggcatctt
                                                                 840
tatttaccaa ctttgcaata acagaaagtc cttccgcctt cgtcgcagag acctcttgct
gggtttcaac aaacgtgcca ttgafricke umgesgatt agacacttcc ccaacgastt
tattgttgag accaagatot gtdaagagta ajaggdaaca gatagagtgt cottggtaat
                                                                 265
aagaagtcag agatttacaa tatgacttta acattaaggt ttatgggata ctcaagatat
                                                                1020
1080
                                                                 1082
<210> 516
<211> 559
<212> DNA
<213> Homo sapiens
<400> 516
ctgctccagc gctgacgccg agccatggcg gacgaggagc ttgaggcgct gaggagacag
                                                                  60
aggetggeeg agetgeagge caaacaeggg gateetggtg atgeggeeca acaggaagea
                                                                 120
aagcacaggg aagcagaaat gagaaacagt atcttagccc aagttctgga tcagtcggcc
                                                                 180
                                                                 240
cgggccaggt taagtaactt agcacttgta aagcctgaaa aaactaaagc agtagagaat
                                                                 300
taccttatac agatggcaag atatggacaa ctaagtgaga aggtatcaga acaaggttta
                                                                 360
atagaaatcc ttaaaaaagt aagccaacaa acagaaaaga caacaacagt gaaattcaac
agaagaaaag taatggactc tgatgaagat gacgattatt gaactacaag tgctcacaga
                                                                 420
ctagaactta acggaacaag tctaggacag aagttaagat ctgattattt actttgttta
                                                                 480
540
aaaaaaaaa aaaaaaaaa
<210> 517
<211> 110
<212> PRT
<213> Homo sapiens
<400> 517
Met Phe Cys Pro Leu Lys Leu Ile Leu Leu Pro Val Leu Leu Asp Tyr
Ser Leu Gly Leu Asn Asp Leu Asn Val Ser Pro Pro Glu Leu Thr Val
                              25
His Val Gly Asp Ser Ala Leu Met Gly Cys Val Phe Gln Ser Thr Glu
```

Asp Lys Cys Ile Phe Lys Ile Asp Trp Thr Leu Ser Pro Gly Glu His 55 Ala Lys Asp Glu Tyr Val Leu Tyr Tyr Tyr Ser Asn Leu Ser Val Pro 70 75 Ile Gly Arg Phe Gln Asn Arg Val His Leu Met Gly Asp Asn Leu Cys 90 Asn Asp Gly Ser Leu Leu Cln Asp Val Gln Asp Val Glu 100 105

<210> 518 <211> 4544 <212> DNA <213> Homo sapiens

<400> 518

ccgagaaggg cttcaggacg cgggaggcgc acttgcttca agtcgcgggc gtgggaacgg 60 ggttgcaaaa cggggccttt ttatccgggc ttgcttccgg cgtcatggct caaagggcct 120 tcccgaatcc ttatgctgat tataacaaat ccctggccga aggctacttt gatgctgccg 180 ggaggctgac tcctgagttc tcacaacgct tgaccaataa gattcgggag cttcttcagc 240 aaatggagag aggcctgaaa tcagcagacc ctcgggatgg caccggttac actggctggg 300 caggtattgc tgtgctttac ttacatcttt atgatgtatt tggggaccct gcctacctac 360 agttagdada tggctatgta aagdarii viiji ihrii haddaagegd tocatcadet 420 teetttgtgg ggatgeagge coeetggway tygwwystyt getatateae aagatgaaca 480 atgagaagca ggcagaagat tgcatcacac ggctaattca cctaaataag attgatcctc atgctccaaa tgaaatgctc tatgggcgaa taggctacat ctatgctctt ctttttgtca ataagaactt tggagtggaa aagatteete aaageeatat teageagatt tgtgaaacaa ttttaacctc tggagaaaac ctagctagga agagaaactt cacggcaaag tctccactga tgtatgaatg gtaccaggaa tattatgtag gggctgctca tggcctggct ggaatttatt actacctgat gcagcccagc cttcaagtga gccaagggaa gttacatagt ttggtcaagc ccagtgtaga ctacgtctgc cagctgaaat tcccttctgg caattaccct ccatgtatag gtgataatcg agatctgctt gtccattggt gccatggcgc ccctggggta atctacatgc tcatccaggc ctataaggta ttcagagagg aaaagtatct ctgtgatgcc tatcagtgtg - 1020 ctgatgtgat ctggcaatat gggttgctga agaagggata tgggctgtgc cacggttctg cagggaatgc ctatgccttc ctgacactct acaacctcac acaggacatg aagtacctgt atagggcctg taagtttgct gaatggtgct tagagtatgg agaacatgga tgcagaacac . cagacaccc tttctctct tttgaaggaa tggctggaac aatatatttc ctggctgacc tgctagtccc cacaaaagcc aggttccctg catttgaact ctgaaaggat agcatgccac ctgcaactca ctgcatgacc ctttctgtat attcaaaccc aagctaagtg cttccgttgc tttccaagga aacaaagagt caaactgtgg acttgatttt gttagctttt ttcagaattt atctttcatt cagttccctt ccattatcat ttacttttac ttagaagtat ccaaggaagt cttttaactt taatttccat ttcttcctaa agggagagtg agtgatatgt acagtgtttt gagattgtat acatatattc cagaacttgg aggaaatctt atttaagttt atgaatataa ccatctgtta ctgttctaaa aatgtttaaa agaaactcaa tacagataaa gataaatatg tgactattat tgggtattac acttcacttc tctttaatat ttttcctcca actggaggge 1740 agacaatttt ctgacttgct tttctctagg tggttcattt tgaaagggga cagaaatata 1800 actaaatgct tccaggagaa aaattccaag agttacaatc tggacttggt acctaaatat 1860 cattttttaa attcttgatg cctatttgga ctagaggtaa acatactttc agattggcct 1920 gtttttgtcg gtaaggcata cagccttcag aagccaacat ttttaatcaa aaacttataa 1980 aacatgatga tcattgtgaa aattctgagt tgaaggttag tttaagataa gctaacaata 2040 acagtotgtg ttttototaa aataatotga gttttttgga actotttatt taaatatgtg tgtttttcag tattcaaata agatcaggaa gccaattttc tatgtatgaa tatgctttaa . cctaggattt cagtccactc tgactgactt tctaaacttt aacttgggtt tttacagtga ctatgcatta gtgctgactc tttggtataa gccataaaat attttccttc ctatcaattt atotgaactt tggtotttto actaaattgt acagtattot acttotqttt aaaaagggga gatgagaaag ggaatactat ctaaccaata acttgaacaa aaacactaaa ctaagcattt aatagaaatg ctttttattg aggaggtatt atccaqactt catgcttaga acaaatgcat ctttgcgtat cctagactta acaattcatc agtttctgag accacagaat caggttttcc gtagtagata aagactetet ggtgetteaa attetgttea agtgttttga eteateaget 2580 tetactettt etattactge etttgeetgg ettgttttgt etetttgeaa etgattttge 2640 aaaaaaaaat tgtagcttta aaataacagg gtctaagtat tttaaatgtg cctatttcac 2700

agctctcttg	gtcacaaaaa	catgctattt	ttattggaac	ttcaaaccaa	atccccactg	2760
agtgtgtact	ggttcctgca	ggtagcagtc	tcctattatc	tcctgtttag	caccaaaaga	2820.
gctaatatta	ttggaaactg	accttttaaa	ggccactggc	agtaggattt	aaaaagcagc	2880
ccactgctca	gtttccagga	tcagcttcct	ccttctgtca	cttgtgtaag	ttggcactac	2940
cttgtgcctc	tcagattgct	gaagtgctgc	tggtaagcat	gtgcatgctc	tgcctttctt	3000
gtgaaagttt	tcaatcagcg	atatcagcac	ttacagtaag	aagtaaaagt	agtgcacagc	3060
aaagctaatt	tgcctttgcc	tggggtgttc	agcttgaaag	aataaagctc	atttggttta	3120
gttaaatgtc	ttactctact	gtgcctatgc	ttttagctgc	gttactaagc	aagggaaaaa	3180
taacaqtttc	tctgagccag	agaagacttg	atcacagttc	tccaagcatc	gtgatagcaa	3240
tgcttaaccc	caggaagatt	tcaaggcagg	gagaagaaca	tttcaaataa	gattcttgtt	3300
aacccattta	tgcctagtgt	tccattattg	gaatgctaag	cttgtgggag	tcatttacat	3360
cctactgctc	aaagtcattg	ccaaqqtctq	atttttcaca	caaaaaattg	caacccccag	3420
cataaatggg	ttagctactg	tcatcagtta	gcaaattcat	ccacacaaac	acaattagag	3480
tttaattttt	ttttaagctt	ttcaaaactt	actaaactgg	cacaatttta	tatgtatgct	3540
atttgttgta	tttatgctta	agagcaaaaa	agttttgatg	ggattttaaa	ttcagcaaag	3600
cctacaacqc	tgagacaatc	ccctaacaac	atggtagtaa	ctaaagaaac	ttttatacta	3660
ggcttcttag	ttttaaaagg	aaqtqqcatc	attqtttcaq	ttctagtttg	tatttttctc	3720
tcagatattt	ttcttcttta	aaaatctttc	ccagaagttg	gttcctagaa	aactcaatac	3780
catcatctct	tatctctata	cagggactag	qtaataaaac	cttcaaaggt	tgtcaaaggt	3840
catcaagcag	tgttcattta	tcctqtcaca	tatttctatt	tctatagtaa	tttagaaatt	3900
gcaaatagtt	aacttttcat	catqtaaaaa	qttaacatta	tcctatttcc	atagatacca	3960
tagacagcag	tgtggcctga	attatcaatc	tttaatcctq	agtcatgtgg	ctctctttc	4020
atctttgatg	tcagttccaa	ttatttggca	tcaaaaacct	tcatqqtaqq	tagagtttta	4080
- 4404	gatctagggt	tactttcl	**	รับเกลเ	'gaattgaga	4340
	gctactatgt	cctcagguta	attices of the	j., Ltauga	geeetgeet	4200
ittactaget	actttagaaa	tagaaaatgt	gaagagtgac	tatttacatg	tatactcctt	4260
tagctactag	aactcatctg	tagtccttta	ttatttacac	tgaattccáa	tttcatttct	4320
cttccqctaa	gtaagagcac	ctcattcctq	tgttttctct	actattgagc	tgtagacgaa	4380
ctatttctct	aattataaag	caaactgttt	gggatattca	gggaaactac	cccaatgtta	4440
	taatgggaaa					4500
	ttctcaataa				-	4544
5-4-1-1-45			JJ	-		

<210> 519

<211> 1779

<212> DNA

<213> Mus musculus

### <400> 519

60 ggtccggaat tcccgggtcg acceacgcgt ccgctggcct tgggcgcaga ccccggccgg 120 tcccggggct gcctctttaa gggaggggt ggagccgcga gtcaggcgcg aggagctcca 180 gaaatcttga ggccagagcc ccgcacctcg gcgcagccat gagtgcggag gtgaaggtga 240 cagggcagaa ccaagagcag tttctgctcc ttgccaagtc ggctaagggg gcggcactgg 300 ccacactcat ccaccaggtg ctggaggccc ctggtgtcta cgtgtttggg gaactgctgg atatgcctaa tgttagagag ctggcagaaa gcgactttgc ctccaccttc cggctgctca 360 cagtgtttgc ctatgggacc tatgcggact acttagctga agccaggaat ctcccccac 420 tgactgacgc acagaagaat aagcttcgac atctgtcagt tgtcactctg gctgccaaag tcaagtgtat cccatatgca gtgttgctgg aggcccttgc ccttcgaaac gtgcgccagc tggaagacct tgtgatcgag gctgtgtatg ctgatgtcct tcgtggctct ctggaccagc 600 660 gcaatcagcg gctagaggtt gattacagca tcgggcggga catccagcgc caggacctca gtgccatcgc ccagaccctg caagagtggt gcgtgggctg tgaggttgtg ttgtcgggca 720 780 tegaagagea ggteageegt geeaaceage acaaggagea geagetggge etgaageage agatcgaaag tgaggttgcc aaccttaaga aaaccattaa agttacgaca gcagctgctg 840 900 ctgcagccac ctcccaggat cctgagcaac acctgacaga gctgagagaa ccagcttctg 960 gcaccaacca gcgccagccc agcaagaaag cctccaaggg caagggactc cgagggagcg ccaagatttg gtccaagtcg aactgaaagg acttgtttct tccctgggaa tgtggggtcc 1020 1080 cagetgeeta cetgeetace cettaggagt ceteagagee tteetgtgee eetggeeage tgataatgct agttcattac ttttcatctc ctccaccccc aagcataagc cacaccctct 1140 1200 gtagggagga ggccagtgca ggtcatgttc tgttggtacc tcttatgtgt tccatgctct 1260 tecceageae gettgetete ategttttte egeaetgtgt etgeceatta cecetgteat 1320 tgagcaggtt ggcagtccta tggagggtgc tggctcttaa ccacccacac ctacccctgc

atocctaato	tacaattcct	cctcctcccc	ttgcctagtg	ggctgcatct	gaaaagccat	1380
acgoodance	gretecacet	tcattccage	cttagagttc	tggagccagt	ctgctaccct	1440
ggggaagggg	gracattttc	ctcccagaac	cccatcacac	tacaattgtt	tctttcctct	1500
gggagccgct	tagacattaga	gatactgctg	cttcagtgac	cccagageet	gagaacagct	1560
CtCatCtCCt	tattaaaaaa	tacttetta	ttactcatca	tottaggaag	cccaatggaa	1620
attttgaga	tyttaayaaa	testestata	attataataa	ggaaggaaat	atagattgta	1680
atcctggaag	gatttatate	teeteetgeg	geeeeggegg	ttaacacata	acacagaaat	1740
				Ligacacacg	acacagaaat	1779
aaatqtatqa	gaaatgtatg	tacaaaaaaa	aaaaaaaa			2.75

## **PCT**

# WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



### INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

	INTERNATIONAL APPLICATION PUBLIS.	HED !	UNDER THE PATENT COOPERATION TREATT (FCT)
[	(51) International Patent Classification ⁶ :		(11) International Publication Number: WO 99/31236
	C12N 15/12, C07K 14/47, 16/18, C12Q 1/68	A3	(43) International Publication Date: 24 June 1999 (24.06.99)
	(21) International Application Number: PCT/IB (22) International Filing Date: 17 December 1998 (		BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR,
ı	(22) International Lang Daves		KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,

US

US

US

US

(71) Applicant (for all designated States except US): GENSET [FR/FR]; 24, rue Royale, F-75008 Paris (FR).

17 December 1997 (17.12.97)

9 February 1998 (09.02.98)

10 August 1998 (10.08.98)

13 April 1998 (13.04.98)

(72) Inventors; and

(30) Priority Data:

60/069,957

60/074,121

60/081,563

60/096,116

- (75) Inventors/Applicants (for US only): BOUGUELERET, Lydie [FR/FR]; 108, avenue Victor Hugo, F-92170 Vanves (FR). DUCLERT, Aymeric [FR/FR]; 6 ter, rue Victorine, F-94100 Saint-Maur (FR). DUMAS MILNE EDWARDS, Jean-Baptiste [FR/FR]; 8, rue Grégoire de Tours, F-75006 Paris (FR).
- (74) Agents: MARTIN, Jean-Jacques et al.; Cabinet Regimbeau, 26, avenue Kléber, F-75116 Paris (FR).

B1) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

#### Published

With international search report.

Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(88) Date of publication of the international search report: 10 September 1999 (10.09.99)

(54) Title: EXTENDED cDNAs FOR SECRETED PROTEINS

(57) Abstract

The sequences of extended cDNAs encoding secreted proteins are disclosed. The extended cDNAs can be used to express secreted proteins or portions thereof or to obtain antibodies capable of specifically binding to the secreted proteins. The extended cDNAs may also be used in diagnostic, forensic, gene therapy, and chromosome mapping procedures. The extended cDNAs may also be used to design expression vectors and secretion vectors.

## FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia.
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
ΑU	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav	TM	Turkmenistan
BF	Burkina Faso	GR	Greece		Republic of Macedonia	TR	Turkey
BG	Bulgaria	HU	Hungary	ML	Mali	TT	Trinidad and Tobago
BJ	Benin	IE	Ireland	MN	Mongolia	UA	Ukraine,
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of Americ
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
СН	Switzerland	KG	Kyrgyzstan	NO	Norway	zw	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's	NZ	New Zealand		
CM	Cameroon		Republic of Korea	PL	Poland		
CN	China	KR	Republic of Korea	PT	Portugal		
CU	Cuba	KZ	Kazakstan	RO	Romania		
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation		
DE	Germany	LI	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		

international Application No
' / IB 98/02122

A. CLASSIFICATION OF SUBJECT MATTER IPC 6 C12N15/12 C07 C07K14/47 C07K16/18 C12Q1/68 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) C12N C07K C12Q Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. E,L WO 99 06549 A (GENSET (FR); DUMAS MILNE 1-20 EDWARDS J.-B.; DUCLERT A.; LACROIX B.) 11 February 1999 (1999-02-11) L: Priority abstract page 6 - page 12
page 129 - page 133; claims Seq. ID:251 page 213 - page 214 Seq.ID:484 page 366 - page 367 Χ Database EMBL, entry HS695112 2,5,8 Accession number R50695 24 May 1995 95% identity with Seq.ID:40 nt.1-384 XP002097725 the whole document -/--Χ Further documents are listed in the continuation of box C. X Patent family members are listed in annex. Special categories of cited documents : "T" later document published after the international filing date or priority date and not in conflict with the application but "A" document defining the general state of the art which is not cited to understand the principle or theory underlying the considered to be of particular relevance "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention filing date cannot be considered novel or cannot be considered to document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the citation or other special reason (as specified) *O* document referring to an oral disclosure, use, exhibition or document is combined with one or more other such docu-ments, such combination being obvious to a person skilled in the art. document published prior to the international filing date but later than the priority date claimed *&* document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report **2** 7. 07. 99 24 March 1999 Name and mailing address of the ISA Authorized officer European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo ni, Macchia, G Fax: (+31-70) 340-3016

F /IB 98/02122

C.(Continua	(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT					
Category 3		Relevant to claim No.				
A	WO 96 34981 A (GENSET (FR); NICOLAEVNA MERENKOVA I.; DUMAS MILNE EDWARDS JB.G.) 7 November 1996 (1996-11-07) cited in the application abstract					
Α	EP 0 625 572 A (KANAGAWA ACAD OF SCIENCE AND TECHNOL FOUNDATION (JP); KATO S; SEKINE S) 23 November 1994 (1994-11-23) cited in the application abstract					
A	CARNINCI P. ET AL.: "High-efficiency full-length cDNA cloning by biotinylated CAP trapper" GENOMICS, vol. 37, no. 3, 1 November 1996 (1996-11-01), pages 327-336, XP002081729 cited in the application abstract					
A	KATO S. ET AL.: "Construction of a human full-length cDNA bank" GENE, vol. 150, 1994, pages 243-250, XP002081364 cited in the application abstract					
A	WO 97 07198 A (GENETICS INSTITUTE INC (US); JACOBS K; MCCOY JM; KELLEHER K; CARLIN M) 27 February 1997 (1997-02-27)					
A	TASHIRO K. ET AL.: "Signal sequence trap: a cloning strategy for secreted proteins and type I membrane proteins" SCIENCE, vol. 261, 30 July 1993 (1993-07-30), pages 600-603, XP000673204 abstract					
Α	YOKOYAMA-KOBAYASHI M. ET AL.: "A signal sequence detection system using secreted protease activity as an indicator" GENE, vol. 163, 1995, pages 193-196, XP002053953 abstract					
A	HEIJNE VON G.: "A new method for predicting signal sequence cleavage sites" NUCLEIC ACIDS RESEARCH, vol. 14, no. 11, 1986, pages 4683-4690, XP002053954 cited in the application abstract					
	-/					

......

International Application No
7/IB 98/02122

		./18 98/02122				
C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT						
Category °	ory ° Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No.					
A	LOCKHART D.J. ET AL.: "Expression monitoring by hybridization to high-density oligonucleotide arrays" BIO/TECHNOLOGY, no. 14, 14 December 1996 (1996-12-14), pages 1675-1680, XP002074420 abstract	18				
	·					

International application No.

PCT/IB 98/02122

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Claims Nos.:  because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
See additional sheet.
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
A. X No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  Invention 1, Claims 1-20 partially.
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

1. Claims: Invention 1: Claims 1-20, all partially.

Nucleic acid comprising the sequence as in Seq.ID:40, complementary sequence or fragments, host cell containing said nucleic acid. Polypeptide as in Seq.ID:141, encoded by said polynucleotide, or fragments, method of making said polypeptide. Antibody specifically binding to said polypeptide.

2. Claims: Inventions 2-233: Claims 1-20, all partially, as far as applicable.

Idem as subject 1 but limited to each of the DNA sequences as in Seq.ID:41-140, 242-377, and corresponding polypeptides, where invention 2 is limited to Seq.ID:41 and 142, invention 3 is limited to Seq.ID:42 and 143, ....., invention 8 is limited to Seq.ID:47 and 148, invention 9 is limited to Seq.ID:48,49,110,149,150 and 211, invention 10 is limited to Seq.ID:50 and 151, ....., invention 32 is limited to Seq.ID:72 and 173, invention 33 is limited to Seq.ID:73,74,131,174,175 and 232, invention 34 is limited to Seq.ID:75 and 176, ....., invention 233 is limited to Seq.ID:377 and 513.

For the sake of conciseness, the first subject matter is explicitly defined, the other subject matters are defined by analogy thereto.

information on paterit family members

International Application No

F /IB 98/02122

Patent document cited in search report		Publication date	Patent family member(s)		Publication date	
WO 9906549	Α	11-02-1999	AU	8555098 A	22-02-1999	
WO 9634981	A	07-11-1996	FR FR AU CA EP	2733765 A 2733762 A 5982996 A 2220045 A 0824598 A	08-11-1996 08-11-1996 21-11-1996 07-11-1996 25-02-1996	
EP 0625572	Α	23-11-1994	JP WO US	6153953 A 9408001 A 5597713 A	03-06-1994 14-04-1994 28-01-1997	
WO 9707198	A	27-02-1997	US AU AU CA CA EP EP WO	5707829 A 6712396 A 6768596 A 2227220 A 2229208 A 0839196 A 0851875 A 9704097 A	13-01-1998 18-02-1997 12-03-1997 06-02-1997 27-02-1997 06-05-1998 08-07-1998 06-02-1997	